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SITIONS THEREOF

(57) Abstract

Recombinant or substantially pure preparations of *H. pylori* polypeptides are described. The nucleic acids encoding the polypeptides also are described. The *H. pylori* polypeptides are useful for diagnostics and vaccine compositions.

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NUCLEIC ACID AND AMINO ACID SEQUENCES RELATING TO HELICOBACTER PYLORI AND VACCINE COMPOSITIONS THEREOF

Background of the Invention

Helicobacter pylori is a gram-negative, S-shaped, microaerophilic bacterium that was discovered and cultured from a human gastric biopsy specimen. (Warren, J.R. and B. Marshall, (1983) Lancet 1: 1273-1275; and Marshall et al., (1984) Microbios Lett. 25: 83-88). H. pylori has been strongly linked to chronic gastritis and duodenal ulcer disease. (Rathbone et. al., (1986) Gut 27: 635-641). Moreover, evidence is accumulating for an etiologic role of H. pylori in nonulcer dyspepsia, gastric ulcer disease, and gastric adenocarcinoma. (Blaser M. J., (1993) Trends Microbiol. 1: 255-260). Transmission of the bacteria occurs via the oral route, and the risk of infection increases with age. (Taylor, D.N. and M. J. Blaser, (1991) Epidemiol. Rev 13: 42-50). H. pylori colonizes the human gastric mucosa, establishing an infection that usually persists for decades. Infection by H. pylori is prevalent worldwide. Developed countries have infection rates over 50% of the adult population, while developing countries have infection rates reaching 90% of the adults over the age of 20. (Hopkins R. J. and J. G. Morris (1994) Am. J. Med. 97: 265-277).

The bacterial factors necessary for colonization of the gastric environment, and for virulence of this pathogen, are poorly understood. Examples of the putative virulence factors include the following: urease, an enzyme that may play a role in neutralizing gastric acid pH (Eaton et al., (1991) *Infect. Immunol.* <u>59</u>: 2470-2475; Ferrero, R.L. and A. Lee (1991) *Microb. Ecol. Hlth. Dis.* <u>4</u>: 121-134; Labigne et al., (1991) *J. Bacteriol.* <u>173</u>: 1920-1931); the bacterial flagellar proteins responsible for motility across the mucous layer. (Hazell et al., (1986) *J. Inf. Dis.* <u>153</u>: 658-663; Leying et al., (1992) *Mol. Microbiol.* <u>6</u>: 2863-2874; and Haas et al., (1993) *Mol. Microbiol.* <u>8</u>: 753-760); Vac A, a bacterial toxin that induces the formation of intracellular vacuoles in epithelial cells (Schmitt, W. and R. Haas, (1994) *Molecular Microbiol.* <u>12(2)</u>: 307-319); and several gastric tissue-specific adhesins. (Boren et al., (1993) *Science* <u>262</u>: 1892-1895; Evans et al., (1993) *J. Bacteriol.* <u>175</u>: 674-683; and Falk et al., (1993) *Proc. Natl. Acad. Sci.* USA <u>90</u>: 2035-203).

Numerous therapeutic agents are currently available that eradicate *H. pylori* infections *in vitro*. (Huesca et. al., (1993) *Zbl. Bakt.* 280: 244-252; Hopkins, R. J. and J. G. Morris, supra). However, many of these treatments are suboptimally effective *in vivo* because of bacterial resistance, altered drug distribution, patient non-compliance or poor drug availabilty. (Hopkins, R. J. and J. G. Morris, supra). Treatment with antibiotics combined with bismuth are part of the standard regime used to treat *H. pylori* infection.

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(Malfertheiner, P. and J. E. Dominguez-Munoz (1993) Clinical Therapeutics 15 Supp. B: 37-48). Recently, combinations of a proton pump inhibitors and a single antibiotic have been shown to ameliorate duodenal ulcer disease. (Malfertheiner, P. and J. E. Dominguez-Munoz supra). However, methods employing antibiotic agents can have the problem of the emergence of bacterial strains which are resistant to these agents. (Hopkins, R. J. and J. G. Morris, supra). These limitations demonstrate that new more effective methods are needed to combat *H. pylori* infections *in vivo*. In particular, the design of new vaccines that may prevent infection by this bacterium is highly desirable.

Summary of the Invention

This invention relates to novel genes, e.g., genes encoding polypeptides such as bacterial surface proteins, from the organism *Helicobacter pylori* (*H. pylori*), and other related genes, their products, and uses thereof. The nucleic acids and peptides of the present invention have utility for diagnostic and therapeutics for *H. pylori* and other *Helicobacter* species. They can also be used to detect the presence of *H. pylori* and other *Helicobacter* species in a sample; and for use in screening compounds for the ability to interfere with the *H. pylori* life cycle or to inhibit *H. pylori* infection. More specifically, this invention features compositions of nucleic acids corresponding to entire coding sequences of *H. pylori* proteins, including surface or secreted proteins or parts thereof, nucleic acids capable of binding mRNA from *H. pylori* proteins to block protein translation, and methods for producing *H. pylori* proteins or parts thereof using peptide synthesis and recombinant DNA techniques. This invention also features antibodies and nucleic acids useful as probes to detect *H. pylori* infection. In addition, vaccine compositions and methods for the protection or treatment of infection by *H. pylori* are within the scope of this invention.

Detailed Description of the Drawings

Figure 1 is a bar graph that depicts the antibody titer in serum of mice following immunization with specific *H. pylori* antigens.

Figure 2 is a bar graph that depicts the antibody titer in mucous of mice following immunization with specific *H. pylori* antigens.

Figure 3 is a bar graph that depicts therapeutic immunization of *H. pylori* infected mice with specific antigens dissolved in HEPES buffer.

Figure 4 is a bar graph that depicts therapeutic immunization of *H. pylori* infected mice with specific antigens dissolved in buffer containing DOC.

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Figure 5 depicts the amino acid sequence alignment in a portion of the sequence of five *H. pylori* proteins (depicted in the single letter amino acid code; shown N-terminal to C-terminal, left to right).

Figure 6 depicts the amino acid sequence alignment in a portion of the sequence of four *H. pylori* proteins (depicted in the single letter amino acid code; shown N-terminal to C-terminal, left to right).

Figure 7 depicts the amino acid sequence alignment in a portion of the sequence of two *H. pylori* proteins (depicted in the single letter amino acid code; shown N-terminal to C-terminal, left to right).

Figure 8 depicts the amino acid sequence alignment in a portion of the sequence of two *H. pylori* proteins (depicted in the single letter amino acid code; shown N-terminal to C-terminal, left to right).

Detailed Description of the Invention

In one aspect, the invention features a recombinant or substantially pure preparation of *H. pylori* polypeptide of SEQ ID NO: 74. The invention also includes substantially pure nucleic acid encoding an *H. pylori* polypeptide of SEQ ID NO: 74, such nucleic acid is contained in SEQ ID NO: 1. The *H. pylori* polypeptide sequences of the invention described herein are contained in the Sequence Listing, and the nucleic acids encoding *H. pylori* polypeptides of the invention are contained in the Sequence Listing.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 75, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 2.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 76, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 3.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 77, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 4.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 78, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 5.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 79, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 6.

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In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 80, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 7.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 81, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 8.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 82, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 9.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 83, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 10.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 84, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 11.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 85, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 12.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 86, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 13.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 87, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 14.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 88, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 15.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 89, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 16.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 90, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 17.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 91, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 18.

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In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 92, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 19.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 93, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 20.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 94, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 21.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 95, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 22.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 96, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 23.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 97, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 24.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 98, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 25.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 99, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 26.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 100, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 27.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 101, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 28.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 102, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 29.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 103, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 30.

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In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 104, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 31.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 105, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 32.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 106, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 33.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 107, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 34.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 108, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 35.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 109, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 36.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 110, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 37.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 111, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 38.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 112, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 39.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 113, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 40.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 114, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 41.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 115, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 42.

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In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 116, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO:43.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 117, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 44.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 118, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 45.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 119, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 46.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 120, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 47.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 121, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 48.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 122, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 49.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 123, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 50.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 124, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 51.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 125, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 52.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 126, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 53.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 127, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 54.

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In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 128, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 55.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 129, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 56.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 130, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 57.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 131, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 58.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 132, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 59.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 133, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 60.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 134, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 61.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 135, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 62.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 136, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 63.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 137, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 64.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 138, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 65.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 139, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 66.

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In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 140, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 67.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 141, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 68.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 142, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 69.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 143, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 70.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 144, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 71.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 145, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 72.

In another aspect, the invention features a substantially pure nucleic acid encoding an *H. pylori* polypeptide having an amino acid sequence of SEQ ID NO: 146, such as a nucleic acid comprising a nucleotide sequence of SEQ ID NO: 73.

Particularly perferred is an isolated nucleic acid comprising a nucleotide sequence encoding an *H. pylori* cell envelope polypeptide or a fragment thereof. Such nucleic acid is selected from the group consisting of SEQ ID NO: 3, SEQ ID NO: 25, SEQ ID NO: 48, SEQ ID NO: 16, SEQ ID NO: 10, SEQ ID NO: 45, SEQ ID NO: 35, SEQ ID NO: 37, SEQ ID NO: 7, SEQ ID NO: 39, SEQ ID NO: 55, SEQ ID NO: 18, SEQ ID NO: 19, SEQ ID NO: 28, SEQ ID NO: 30, SEQ ID NO: 52, SEQ ID NO: 54, SEQ ID NO: 56, SEQ ID NO: 58, SEQ ID NO: 1, SEQ ID NO: 42, SEQ ID NO: 14, SEQ ID NO: 43, SEQ ID NO: 11, SEQ ID NO: 71, SEQ ID NO: 17, SEQ ID NO: 57, SEQ ID NO: 5, SEQ ID NO: 6, SEQ ID NO: 8, and SEQ ID NO: 21.

In another embodiment, the *H. pylori* cell envelope polypeptide or a fragment thereof is an *H. pylori* inner membrane polypeptide or a fragment thereof encoded by the nucleic acid selected from the group consisting of SEQ ID NO: 3, SEQ ID NO: 25, and SEQ ID NO: 48.

In another embodiment, the *H. pylori* cell envelope polypeptide or a fragment thereof is an *H. pylori* outer membrane polypeptide or a fragment thereof encoded by the nucleic acid selected from the group consisting of SEQ ID NO: 16, SEQ ID NO: 10,

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SEQ ID NO: 45, SEQ ID NO: 35, SEQ ID NO: 37, SEQ ID NO: 7, SEQ ID NO: 39, SEQ ID NO: 55, SEQ ID NO: 18, SEQ ID NO: 19, SEQ ID NO: 28, SEQ ID NO: 30, SEQ ID NO: 52, SEQ ID NO: 54, SEQ ID NO: 56, SEQ ID NO: 58, SEQ ID NO: 1, SEQ ID NO: 42, SEQ ID NO: 14, SEQ ID NO: 43, SEQ ID NO: 11, and SEQ ID NO: 71.

In another embodiment, the *H. pylori* outer membrane polypeptide or a fragment thereof is an *H. pylori* polypeptide having a terminal phenylalanine residue and a C-terminal tyrosine cluster or a fragment thereof encoded by the nucleic acid selected from the group consisting of SEQ ID NO: 1, SEQ ID NO: 42, SEQ ID NO: 14, SEQ ID NO: 43, SEQ ID NO: 11, and SEQ ID NO:71.

In yet another embodiment, the *H. pylori* outer membrane polypeptide or a fragment thereof is an *H. pylori* polypeptide having a terminal phenylalanine residue or a fragment thereof encoded by the nucleic acid selected from the group consisting of SEQ ID NO: 16, SEQ ID NO: 45, SEQ ID NO: 35, SEQ ID NO: 37, SEQ ID NO: 7, SEQ ID NO: 39, SEQ ID NO: 55, SEQ ID NO: 18, SEQ ID NO: 19, SEQ ID NO: 28, SEQ ID NO: 30, SEQ ID NO: 52, SEQ ID NO: 54, SEQ ID NO: 56, and SEQ ID NO: 58.

Particularly preferred is an isolated nucleic acid comprising a nucleotide sequence encoding an *H. pylori* secreted polypeptide or a fragment thereof. Such nucleic acid is selected from the group consisting of SEQ ID NO: 72, SEQ ID NO: 32, SEQ ID NO: 51, SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 9, SEQ ID NO: 13, SEQ ID NO: 22, SEQ ID NO: 29, SEQ ID NO: 31, SEQ ID NO: 33, SEQ ID NO: 34, SEQ ID NO: 36, SEQ ID NO: 38, SEQ ID NO: 40. SEQ ID NO: 41, SEQ ID NO: 44, SEQ ID NO: 46, SEQ ID NO: 49, SEQ ID NO: 53, SEQ ID NO: 59, SEQ ID NO: 61, SEQ ID NO: 62, SEQ ID NO: 63, SEQ ID NO: 65, SEQ ID NO: 66, SEQ ID NO: 67, and SEQ ID NO: 68.

Particularly preferred is an isolated nucleic acid comprising a nucleotide sequence encoding an *H. pylori* cellular polypeptide or a fragment thereof. Such nucleic acid is selected from the group consisting of SEQ ID NO: 12, SEQ ID NO: 15, SEQ ID NO: 20, SEQ ID NO: 23, SEQ ID NO: 24, SEQ ID NO: 26, SEQ ID NO: 27, SEQ ID NO: 47, SEQ ID NO: 50, SEQ ID NO: 60, SEQ ID NO: 64, SEQ ID NO: 69, SEQ ID NO: 70, and SEQ ID NO: 73.

Particularly preferred is a purified or isolated *H. pylori* cell envelope polypeptide or a fragment thereof, wherein the polypeptide is selected from the group consisting of SEQ ID NO: 76, SEQ ID NO: 98, SEQ ID NO: 121, SEQ ID NO: 89, SEQ ID NO: 83, SEQ ID NO: 118, SEQ ID NO: 108, SEQ ID NO: 110, SEQ ID NO: 80, SEQ ID NO: 112, SEQ ID NO: 128, SEQ ID NO: 91, SEQ ID NO: 92, SEQ ID NO: 101, SEQ ID

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NO: 103, SEQ ID NO: 125, SEQ ID NO: 127, SEQ ID NO: 129, SEQ ID NO: 131, SEQ ID NO: 74, SEQ ID NO: 115, SEQ ID NO: 87, SEQ ID NO: 116, SEQ ID NO: 84, SEQ ID NO: 144, SEQ ID NO: 90, SEQ ID NO: 130, SEQ ID NO: 78, SEQ ID NO: 79, SEQ ID NO: 81, and SEQ ID NO: 94.

In another embodiment, the *H. pylori* cell envelope polypeptide or a fragment thereof is an *H. pylori* inner membrane polypeptide or a fragment thereof selected from the group consisting of SEQ ID NO: 76, SEQ ID NO: 98, and SEQ ID NO: 121.

In another embodiment, the *H. pylori* cell envelope polypeptide or a fragment thereof is an *H. pylori* outer membrane polypeptide or a fragment thereof selected from the group consisting of SEQ ID NO: 89, SEQ ID NO: 83, SEQ ID NO: 118, SEQ ID NO: 108, SEQ ID NO: 110, SEQ ID NO: 80, SEQ ID NO: 112, SEQ ID NO: 128, SEQ ID NO: 91, SEQ ID NO: 92, SEQ ID NO: 101, SEQ ID NO: 103, SEQ ID NO: 125, SEQ ID NO: 127, SEQ ID NO: 129, SEQ ID NO: 131, SEQ ID NO: 74, SEQ ID NO: 115, SEQ ID NO: 87, SEQ ID NO: 116, SEQ ID NO: 84, SEQ ID NO: 144, SEQ ID NO: 90, and SEQ ID NO: 130.

In another embodiment, the *H. pylori* outer membrane polypeptide or a fragment thereof is an *H. pylori* polypeptide having a terminal phenylalanine residue and a C-terminal tyrosine cluster or a fragment thereof selected from the group consisting of SEQ ID NO: 74, SEQ ID NO: 115, SEQ ID NO: 87, SEQ ID NO: 116, SEQ ID NO: 84, and SEQ ID NO:144.

In another embodiment, the *H. pylori* outer membrane polypeptide or a fragment thereof is an *H. pylori* polypeptide having a terminal phenylalanine residue or a fragment thereof selected from the group consisting of SEQ ID NO: 89, SEQ ID NO: 118, SEQ ID NO: 108, SEQ ID NO: 110, SEQ ID NO: 80, SEQ ID NO: 112, SEQ ID NO: 128, SEQ ID NO: 91, SEQ ID NO: 92, SEQ ID NO: 101, SEQ ID NO: 103, SEQ ID NO: 125, SEQ ID NO: 127, SEQ ID NO: 129, SEQ ID NO: 131.

Particularly preferred is a purified or isolated *H. pylori* secreted polypeptide or a fragment thereof, wherein the polypeptide is selected from the group consisting of SEQ ID NO: 145, SEQ ID NO: 105, SEQ ID NO: 124, SEQ ID NO: 75, SEQ ID NO: 77, SEQ ID NO: 82, SEQ ID NO: 86, SEQ ID NO: 95, SEQ ID NO: 102, SEQ ID NO: 104, SEQ ID NO: 106, SEQ ID NO: 107, SEQ ID NO: 109, SEQ ID NO: 111, SEQ ID NO: 113, SEQ ID NO: 114, SEQ ID NO: 117, SEQ ID NO: 119, SEQ ID NO: 122, SEQ ID NO: 126, SEQ ID NO: 132, SEQ ID NO: 134, SEQ ID NO: 135, SEQ ID NO: 136, SEQ ID NO: 138, SEQ ID NO: 139, SEQ ID NO: 140, and SEQ ID NO: 141.

Particularly preferred is a purified or isolated *H. pylori* cellular polypeptide or a fragment thereof, wherein the polypeptide is selected from the group consisting of SEQ ID NO: 85, SEQ ID NO: 88, SEQ ID NO: 93, SEQ ID NO: 96, SEQ ID NO: 97, SEQ

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ID NO: 99, SEQ ID NO: 100, SEQ ID NO: 120, SEQ ID NO: 123, SEQ ID NO: 133, SEQ ID NO: 137, SEQ ID NO: 142, SEQ ID NO: 143, and SEQ ID NO: 146.

In another aspect, the invention pertains to any individual *H. pylori* polypeptide member or nucleic acid encoding such a member from the above-identified groups of *H. pylori* polypeptides.

In another aspect, the invention features nucleic acids capable of binding mRNA of *H. pylori*. Such nucleic acid is capable of acting as antisense nucleic acid to control the translation of mRNA of *H. pylori*. A further aspect features a nucleic acid which is capable of binding specifically to an *H. pylori* nucleic acid. These nucleic acids are also referred to herein as complements and have utility as probes and as capture reagents.

In another aspect, the invention features an expression system comprising an open reading frame corresponding to *H. pylori* nucleic acid. The nucleic acid further comprises a control sequence compatible with an intended host. The expression system is useful for making polypeptides corresponding to *H. pylori* nucleic acid.

In another aspect, the invention features a cell transformed with the expression system to produce *H. pylori* polypeptides.

In another aspect, the invention features a method of generating antibodies against *H. pylori* polypeptides which are capable of binding specifically to *H. pylori* polypeptides. Such antibodies have utility as reagents for immunoassays to evaluate the abundance and distribution of *H. pylori*-specific antigens.

In another aspect, the invention features a method of generating vaccines for immunizing an individual against *H. pylori*. The vaccination method includes: immunizing a subject with at least one *H. pylori* polypeptide according to the present invention, e.g., a surface or secreted polypeptide, or active portion thereof, and a pharmaceutically acceptable carrier. Such vaccines have therapeutic and/or prophylactic utilities.

In another aspect, the invention provides a method for generating a vaccine comprising a modified immunogenic *H. pylori* polypeptide, e.g., a surface or secreted polypeptide, or active portion thereof, and a pharmacologically acceptable carrier.

In another aspect, the invention features a method of evaluating a compound, e.g. a polypeptide, e.g., a fragment of a host cell polypeptide, for the ability to bind an H. pylori polypeptide. The method includes: contacting the candidate compound with an H. pylori polypeptide and determining if the compound binds or otherwise interacts with an H. pylori polypeptide. Compounds which bind H. pylori are candidates as activators or inhibitors of the bacterial life cycle. These assays can be performed in vitro or in vivo.

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In another aspect, the invention features a method of evaluating a compound, e.g. a polypeptide, e.g., a fragment of a host cell polypeptide, for the ability to bind an *H. pylori* nucleic acid, e.g., DNA or RNA. The method includes: contacting the candidate compound with an *H. pylori* nucleic acid and determining if the compound binds or otherwise interacts with an *H. pylori* polypeptide. Compounds which bind *H. pylori* are candidates as activators or inhibitors of the bacterial life cycle. These assays can be performed *in vitro* or *in vivo*.

The invention features H. pylori polypeptides, preferably a substantially pure preparation of an H. pylori polypeptide, or a recombinant H. pylori polypeptide. In preferred embodiments: the polypeptide has biological activity; the polypeptide has an amino-acid-sequence at least 60%, 70%, 80%, 90%, 95%, 98%, or 99% identical or homologous to an amino acid sequence of the invention contained in the Sequence Listing, preferably it has about 65% sequence identity with an amino acid sequence of the invention contained in the Sequence Listing, and most preferably it has about 92% to about 99% sequence identity with an amino acid sequence of the invention contained in the Sequence Listing; the polypeptide has an amino acid sequence essentially the same as an amino acid sequence of the invention contained in the Sequence Listing; the polypeptide is at least 5, 10, 20, 50, 100, or 150 amino acid residues in length; the polypeptide includes at least 5, preferably at least 10, more preferably at least 20, more preferably at least 50, 100, or 150 contiguous amino acid residues of the invention contained in the Sequence Listing. In yet another preferred embodiment, the amino acid sequence which differs in sequence identity by about 7% to about 8% from the H. pylori amino acid sequences of the invention contained in the Sequence Listing is also encompassed by the invention.

In preferred embodiments: the *H. pylori* polypeptide is encoded by a nucleic acid of the invention contained in the Sequence Listing, or by a nucleic acid having at least 60%, 70%, 80%, 90%, 95%, 98%, or 99% homology with a nucleic acid of the invention contained in the Sequence Listing.

In a preferred embodiment, the subject *H. pylori* polypeptide differs in amino acid sequence at 1, 2, 3, 5, 10 or more residues from a sequence of the invention contained in the Sequence Listing. The differences, however, are such that the *H. pylori* polypeptide exhibits an *H. pylori* biological activity, e.g., the *H. pylori* polypeptide retains a biological activity of a naturally occurring *H. pylori* polypeptide.

In preferred embodiments, the polypeptide includes all or a fragment of an amino acid sequence of the invention contained in the Sequence Listing; fused, in reading frame, to additional amino acid residues, preferably to residues encoded by genomic

DNA 5' or 3' to the genomic DNA which encodes a sequence of the invention contained in the Sequence Listing.

In yet other preferred embodiments, the *H. pylori* polypeptide is a recombinant fusion protein having a first *H. pylori* polypeptide portion and a second polypeptide portion, e.g., a second polypeptide portion having an amino acid sequence unrelated to *H. pylori*. The second polypeptide portion can be, e.g., any of glutathione-S-transferase, a DNA binding domain, or a polymerase activating domain. In preferred embodiment the fusion protein can be used in a two-hybrid assay.

Polypeptides of the invention include those which arise as a result of alternative transcription events, alternative RNA splicing events, and alternative translational and postranslational events.

The invention also encompasses an immunogenic component which includes at least one *H. pylori* polypeptide in an immunogenic preparation; the immunogenic component being capable of eliciting an immune response specific for the *H. pylori* polypeptide, e.g., a humoral response, an antibody response, or a cellular response. In preferred embodiments, the immunogenic component comprises at least one antigenic determinant from a polypeptide of the invention contained in the Sequence Listing.

In another aspect, the invention provides a substantially pure nucleic acid having a nucleotide sequence which encodes an *H. pylori* polypeptide. In preferred embodiments: the encoded polypeptide has biological activity; the encoded polypeptide has an amino acid sequence at least 60%, 70%, 80%, 90%, 95%, 98%, or 99% homologous to an amino acid sequence of the invention contained in the Sequence Listing; the encoded polypeptide has an amino acid sequence essentially the same as an amino acid sequence of the invention contained in the Sequence Listing; the encoded polypeptide is at least 5, 10, 20, 50, 100, or 150 amino acids in length; the encoded polypeptide comprises at least 5, preferably at least 10, more preferably at least 20, more preferably at least 50, 100, or 150 contiguous amino acids of the invention contained in the Sequence Listing.

In preferred embodiments: the nucleic acid of the invention is that contained in the Sequence Listing; the nucleic acid is at least 60%, 70%, 80%, 90%, 95%, 98%, or 99% homologous with a nucleic acid sequence of the invention contained in the Sequence Listing.

In a preferred embodiment, the encoded *H. pylori* polypeptide differs (e.g., by amino acid substitution, addition or deletion of at least one amino acid residue) in amino acid sequence at 1, 2, 3, 5, 10 or more residues, from a sequence of the invention contained in the Sequence Listing. The differences, however, are such that: the *H*.

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pylori encoded polypeptide exhibits a H. pylori biological activity, e.g., the encoded H. pylori enzyme retains a biological activity of a naturally occurring H. pylori.

In preferred embodiments, the encoded polypeptide includes all or a fragment of an amino acid sequence of the invention contained in the Sequence Listing; fused, in reading frame, to additional amino acid residues, preferably to residues encoded by genomic DNA 5' or 3' to the genomic DNA which encodes a sequence of the invention contained in the Sequence Listing.

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In preferred embodiments, the subject *H. pylori* nucleic acid will include a transcriptional regulatory sequence, e.g. at least one of a transcriptional promoter or transcriptional enhancer sequence, operably linked to the *H. pylori* gene sequence, e.g., to render the *H. pylori* gene sequence suitable for expression in a recombinant host cell.

In yet a further preferred embodiment, the nucleic acid which encodes an *H. pylori* polypeptide of the invention, hybridizes under stringent conditions to a nucleic acid probe corresponding to at least 8 consecutive nucleotides of the invention contained in the Sequence Listing; more preferably to at least 12 consecutive nucleotides of the invention contained in the Sequence Listing; more preferably to at least 20 consecutive nucleotides of the invention contained in the Sequence Listing; more preferably to at least 40 consecutive nucleotides of the invention contained in the Sequence Listing.

In a preferred embodiment, the nucleic acid encodes a peptide which differs by at least one amino acid residue from the sequences of the invention contained in the Sequence Listing.

In a preferred embodiment, the nucleic acid differs by at least one nucleotide from a nucleotide sequence of the invention contained in the Sequence Listing which encodes amino acids of the invention contained in the Sequence Listing.

In another aspect, the invention encompasses: a vector including a nucleic acid which encodes an *H. pylori* polypeptide or an *H. pylori* polypeptide variant as described herein; a host cell transfected with the vector; and a method of producing a recombinant *H. pylori* polypeptide or *H. pylori* polypeptide variant; including culturing the cell, e.g., in a cell culture medium, and isolating the *H. pylori* or *H. pylori* polypeptide variant, e.g., from the cell or from the cell culture medium.

In another aspect, the invention features, a purified recombinant nucleic acid having at least 50%, 60%, 70%, 80%, 90%, 95%, 98%, or 99% homology with a sequence of the invention contained in the Sequence Listing.

The invention also provides a probe or primer which includes a substantially purified oligonucleotide. The oligonucleotide includes a region of nucleotide sequence which hybridizes under stringent conditions to at least 8 consecutive nucleotides of sense or antisense sequence of the invention contained in the Sequence Listing, or

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naturally occurring mutants thereof. In preferred embodiments, the probe or primer further includes a label group attached thereto. The label group can be, e.g., a radioisotope, a fluorescent compound, an enzyme, and/or an enzyme co-factor. Preferably the oligonucleotide is at least 8 and less than 10, 20, 30, 50, 100, or 150 nucleotides in length.

The invention also provides an isolated *H. pylori* polypeptide which is encoded by a nucleic acid which hybridizes under stringent hybridization conditions to a nucleic acid contained in the Sequence Listing.

The invention further provides nucleic acids, e.g., RNA or DNA, encoding a polypeptide of the invention. This includes double stranded nucleic acids as well as coding and antisense single strands.

The *H. pylori* strain, from which genomic sequences have been sequenced, has been deposited in the American Type Culture Collection (ATCC # 55679; deposited by Genome Therapeutics Corporation, 100 Beaver Street, Waltham, MA 02154) as strain HP-J99.

Included in the invention are: allelic variations; natural mutants; induced mutants; proteins encoded by DNA that hybridizes under high or low stringency conditions to a nucleic acid which encodes a polypeptide of the invention contained in the Sequence Listing (for definitions of high and low stringency see Current Protocols in Molecular Biology, John Wiley & Sons, New York, 1989, 6.3.1 - 6.3.6 and 6.4.1-6.4.10, hereby incorporated by reference); and, polypeptides specifically bound by antisera to *H. pylori* polypeptides, especially by antisera to an active site or binding domain of *H. pylori* polypeptide. The invention also includes fragments. preferably biologically active fragments. These and other polypeptides are also referred to herein as *H. pylori* polypeptide analogs or variants.

Putative functions have been determined for several of the *H. pylori* polypeptides of the invention, as shown in Table 1.

Accordingly, uses of the claimed *H. pylori* polypeptides based on these identified functions, as well as other functions as described herein, are also within the scope of the invention.

In addition, the present invention encompasses *H. pylori* polypeptides characterized as shown in Table 1 below, including: *H. pylori* cell envelope proteins, *H. pylori* secreted proteins, and *H. pylori* cellular proteins. Members of these groups were identified by BLAST homology searches and by searches for secretion signal or transmembrane protein motifs. Polypeptides related by significant homology to the polypeptides of Table 1 are also considered to be classified in the manner of the homologs shown in Table 1.

TABLE 1

ORF Name and Group	nt SeqID	aa SeqID
A. CELL ENVELOPE		,
A.1 Inner membrane proteins		
02ge11622_23494043_f1_6	3	76
hp5p15212_13095752_c3_36	25	98
06ep30223_20173437_f1_37	48	121
A.2 Outer membrane proteins	+	12-1
05ee10816 14495437 f2 13	10	83
A.2.1 Terminal phe residue	 	
06ep11509_35954752_f2_1	16	89
06ep10615_14495437_f3_47	45	
03ae10804_14495437_i3_47	35	108
05ae30220_917200_c3_172	37	110
04cp11202_23646885_f2_26	7	80
05ep10815_16131925_c2_97	39	112
09cp61003_5860877_f2_23	55	128
09ae10512_48768_c3_67	18	91
09cp11003_5860877_f3_7	19	92
hp6e12267_30478562_f3_33	28	101
06cp30603_34174212_c3_71	30	103
09cp10224_1962590_f3_31	52	125
09cp61003_30478562_c3_106	54	127
11ae80818 10553192_f2_16	56	129
11ee11408 10584582_c3_51	58	131
A.2.2 Terminal phe residue and C-	- 36	131
terminal tyrosine cluster		
01ae12001_116018_c2_40	1	74
06ap10609_116018_c3_50	42	115
06cp30603_4687507_f1_9	14	87
06cp30603_4687507_f1_7	43	116
05ee10816_36126938_f3_16	11	84
01cp20708 4960952 c1 43	71	144
A.3 Via homolgy		
07ap80601_5083193_f3_8	17	90
11ap20714_4797137_f3_45	57	
A.4 Other cell envelope proteins		
04ap12016_25501501_f1_1	5	78
04cp11202_20415937_f2_25	6	79
04ee11108_3906963_f1_7	8	81
29ep10720_25501501_c2_33	21	94
B. SECRETED PROTEINS	 	
hp3e10342_22448587_c2_15	72	145
hp5p15212_24276587_f1_2	32	105
09ce10413_35336707_f2_9	51	124
0300 104 10_00000101 _12_0		74-7

01ae12001_32462543_c2_43	2	75
03ee11215_1416312_c3_35	4	77
05ae30220_14570443_c2_94	9	82
06cp30603_2772578_c1_46	13	86
29ep10720_289077_f2_12	22	95
03ee11215_22542803_f1_7	29	102
09ae10512_3166040_c1_40	31	104
01ce11104_10742963_c2_12	33	106
02ge10116_36335436_f3_66	34	107
04ep41903_11876461_f1_4	36	109
05ce10208_23631292_f1_6	38	111
05ep10815_22447252_c3_110	40	113
05ep10815_30283516_c3_109	41	114
06ee30709_33851038_c3_30	44	117
06ep11202_21687842_c3_35	46	119
06ep30223_2774062_f1_33	49	122
09cp10713_23912707_c1_26	53	126
11ee11408_4882318_f3_24	59	132
hp4e13394_5908553_f1_1	61	134
hp4e53394_1416312_c3_119	62	135
hp5e15211_24328910_c3_38	63	136
hp6p10606_23493756_c1_21	65	138
hp6p22217_23564012_f1_5	66	139
hp6p22217_272058_f1_2	67	140
hp6p22217_2922143_f2_9	68	141
C. OTHER CELLULAR PROTEINS		
06ap11119_14726542_f3_21	12	85
06ee10709_6136430_c1_11	15	88
12ap10605_14094816_c1_5	20	93
hp2p10272_34042518_f1_2	23	96
hp5e15211_25411557_c1_22	24	97
hp5p15641_3907968_f1_3	26	99
hp6e10967_657638_f3_9	27	100
06ep11202_4569693_c2_28	47	120
06ep30223_3930468_c1_110	50	123
hp2e10911_960952_c2_86	60	133
hp6p10509_14642217_c2_17	64	137
hp6p80503_20964382_f2_11	69	142
hp7e10192_5917593_f1_2	70	143
hp6p10509_14642217_c3_25	73	146

[In Table 1, "nt" represents nucleotide Seq. ID number and "aa" represents amino acid Seq. ID number]

5 <u>Definitions</u>

The terms "purified polypeptide" and "isolated polypeptide" and "a substantially pure preparation of a polypeptide" are used interchangeably herein and, as used herein,

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from other proteins, lipids, and nucleic acids with which it naturally occurs. Preferably, the polypeptide is also separated from substances, e.g., antibodies or gel matrix, e.g., polyacrylamide, which are used to purify it. Preferably, the polypeptide constitutes at least 10, 20, 50 70, 80 or 95% dry weight of the purified preparation. Preferably, the preparation contains: sufficient polypeptide to allow protein sequencing; at least 1, 10, or 100 µg of the polypeptide; at least 1, 10, or 100 mg of the polypeptide. Furthermore, the terms "purified polypeptide" and "isolated polypeptide" and "a substantially pure preparation of a polypeptide," as used herein, refer to both a polypeptide obtained from nature or produced by recombinant DNA techniques as described herein.

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For example, an "isolated" or "purified" protein or biologically active portion thereof is substantially free of cellular material or other contaminating proteins from the cell or tissue source from which the H. pylori protein is derived, or substantially free from chemical precursors or other chemicals when chemically synthesized. The language "substantially free of cellular material" includes preparations of H. pylori protein in which the protein is separated from cellular components of the cells from which it is isolated or recombinantly produced. In one embodiment, the language "substantially free of cellular material" includes preparations of H. pylori protein having less than about 30% (by dry weight) of non-H. pylori protein (also referred to herein as a "contaminating protein"), more preferably less than about 20% of non-H. pylori protein, still more preferably less than about 10% of non-H. pylori protein, and most preferably less than about 5% non-H. pylori protein. When the H. pylori protein or biologically active portion thereof is recombinantly produced, it is also preferably substantially free of culture medium, i.e., culture medium represents less than about 20%, more preferably less than about 10%, and most preferably less than about 5% of the volume of the protein preparation.

The language "substantially free of chemical precursors or other chemicals" includes preparations of *H. pylori* protein in which the protein is separated from chemical precusors or other chemicals which are involved in the synthesis of the protein. In one embodiment, the language "substantially free of chemical precursors or other chemicals" includes preparations of *H. pylori* protein having less than about 30% (by dry weight) of chemical precursors or non-*H. pylori* chemicals, more preferably less than about 20% chemical precursors or non-*H. pylori* chemicals, still more preferably less than about 10% chemical precursors or non-*H. pylori* chemicals, and most preferably less than about 5% chemical precursors or non-*H. pylori* chemicals.

A purified preparation of cells refers to, in the case of plant or animal cells, an *in* vitro preparation of cells and not an entire intact plant or animal. In the case of cultured

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cells or microbial cells, it consists of a preparation of at least 10% and more preferably 50% of the subject cells.

A purified or isolated or a substantially pure nucleic acid, e.g., a substantially pure DNA, (are terms used interchangeably herein) is a nucleic acid which is one or both of the following: not immediately contiguous with both of the coding sequences with which it is immediately contiguous (i.e., one at the 5' end and one at the 3' end) in the naturally-occurring genome of the organism from which the nucleic acid is derived; or which is substantially free of a nucleic acid with which it occurs in the organism from which the nucleic acid is derived. The term includes, for example, a recombinant DNA which is incorporated into a vector, e.g., into an autonomously replicating plasmid or virus, or into the genomic DNA of a prokaryote or eukaryote, or which exists as a separate molecule (e.g., a cDNA or a genomic DNA fragment produced by PCR or restriction endonuclease treatment) independent of other DNA sequences. Substantially pure DNA also includes a recombinant DNA which is part of a hybrid gene encoding additional *H. pylori* DNA sequence.

A "contig" as used herein is a nucleic acid representing a continuous stretch of genomic sequence of an organism.

An "open reading frame", also referred to herein as ORF, is a region of nucleic acid which encodes a polypeptide. This region may represent a portion of a coding sequence or a total sequence and can be determined from a stop to stop codon or from a start to stop codon.

As used herein, a "coding sequence" is a nucleic acid which is transcribed into messenger RNA and/or translated into a polypeptide when placed under the control of appropriate regulatory sequences. The boundaries of the coding sequence are determined by a translation start codon at the five prime terminus and a translation stop code at the three prime terminus. A coding sequence can include but is not limited to messenger RNA, synthetic DNA, and recombinant nucleic acid sequences.

A "complement" of a nucleic acid as used herein referes to an anti-parallel or antisense sequence that participates in Watson-Crick base-pairing with the original sequence.

A "gene product" is a protein or structural RNA which is specifically encoded by a gene.

As used herein, the term "probe" refers to a nucleic acid, peptide or other chemical entity which specifically binds to a molecule of interest. Probes are often associated with or capable of associating with a label. A label is a chemical moiety capable of detection. Typical labels comprise dyes, radioisotopes, luminescent and chemiluminescent moieties, fluorophores, enzymes, precipitating agents, amplification

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sequences, and the like. Similarly, a nucleic acid, peptide or other chemical entity which specifically binds to a molecule of interest and immobilizes such molecule is referred herein as a "capture ligand". Capture ligands are typically associated with or capable of associating with a support such as nitro-cellulose, glass, nylon membranes, beads, particles and the like. The specificity of hybridization is dependent on conditions such as the base pair composition of the nucleotides, and the temperature and salt concentration of the reaction. These conditions are readily discernable to one of ordinary skill in the art using routine experimentation.

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Homologous refers to the sequence similarity or sequence identity between two polypeptides or between two nucleic acid molecules. When a position in both of the two compared sequences is occupied by the same base or amino acid monomer subunit, e.g., if a position in each of two DNA molecules is occupied by adenine, then the molecules are homologous at that position. The percent of homology between two sequences is a function of the number of matching or homologous positions shared by the two sequences divided by the number of positions compared x 100. For example, if 6 of 10 of the positions in two sequences are matched or homologous then the two sequences are 60% homologous. By way of example, the DNA sequences ATTGCC and TATGGC share 50% homology. Generally, a comparison is made when two sequences are aligned to give maximum homology.

Nucleic acids are hybridizable to each other when at least one strand of a nucleic acid can anneal to the other nucleic acid under defined stringency conditions.

Stringency of hybridization is determined by: (a) the temperature at which hybridization and/or washing is performed; and (b) the ionic strength and polarity of the hybridization and washing solutions. Hybridization requires that the two nucleic acids contain complementary sequences; depending on the stringency of hybridization, however, mismatches may be tolerated. Typically, hybridization of two sequences at high stingency (such as, for example, in a solution of 0.5X SSC, at 65° C) requires that the sequences be essentially completely homologous. Conditions of intermediate stringency (such as, for example, 2X SSC at 65° C) and low stringency (such as, for example 2X SSC at 55° C), require correspondingly less overall complementarity between the hybridizing sequences. (1X SSC is 0.15 M NaCl, 0.015 M Na citrate). A preferred, non-limiting example of stringent hybridization conditions are hybridization in 6X sodium chloride/sodium citrate (SSC) at about 45°C, followed by one or more washes in 0.2 X SSC, 0.1% SDS at 50-65°C.

The terms peptides, proteins, and polypeptides are used interchangeably herein.

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As used herein, the term "surface protein" refers to all surface accessible proteins, e.g. inner and outer membrane proteins, proteins adhering to the cell wall, and secreted proteins.

A polypeptide has *H. pylori* biological activity if it has one, two and preferably more of the following properties: (1) if when expressed in the course of an *H. pylori* infection, it can promote, or mediate the attachment of *H. pylori* to a cell; (2) it has an enzymatic activity, structural or regulatory function characteristic of an *H. pylori* protein; (3) the gene which encodes it can rescue a lethal mutation in an *H. pylori* gene; (4) or it is immunogenic in a subject. A polypeptide has biological activity if it is an antagonist, agonist, or super-agonist of a polypeptide having one of the above-listed properties.

A biologically active fragment or analog is one having an *in vivo* or *in vitro* activity which is characteristic of the *H. pylori* polypeptides of the invention contained in the Sequence Listing, or of other naturally occurring *H. pylori* polypeptides, e.g., one or more of the biological activities described herein. Especially preferred are fragments which exist *in vivo*, e.g., fragments which arise from post transcriptional processing or which arise from translation of alternatively spliced RNA's. Fragments include those expressed in native or endogenous cells as well as those made in expression systems, e.g., in CHO cells. Because peptides such as *H. pylori* polypeptides often exhibit a range of physiological properties and because such properties may be attributable to different portions of the molecule, a useful *H. pylori* fragment or *H. pylori* analog is one which exhibits a biological activity in any biological assay for *H. pylori* activity. Most preferably the fragment or analog possesses 10%, preferably 40%, more preferably 60%, 70%, 80% or 90% or greater of the activity of *H. pylori*, in any *in vivo* or *in vitro* assay.

Analogs can differ from naturally occurring *H. pylori* polypeptides in amino acid sequence or in ways that do not involve sequence, or both. Non-sequence modifications include changes in acetylation, methylation, phosphorylation, carboxylation, or glycosylation. Preferred analogs include *H. pylori* polypeptides (or biologically active fragments thereof) whose sequences differ from the wild-type sequence by one or more conservative amino acid substitutions or by one or more non-conservative amino acid substitutions, deletions, or insertions which do not substantially diminish the biological activity of the *H. pylori* polypeptide. Conservative substitutions typically include the substitution of one amino acid for another with similar characteristics, e.g., substitutions within the following groups: valine, glycine; glycine, alanine; valine, isoleucine, leucine; aspartic acid, glutamic acid; asparagine, glutamine; serine, threonine; lysine, arginine; and phenylalanine, tyrosine. Other conservative substitutions can be made in view of the table below.

TABLE 2
CONSERVATIVE AMINO ACID REPLACEMENTS

For Amino Acid	Code	Replace with any of
Alanine	A	D-Ala, Gly, beta-Ala, L-Cys, D-Cys
Arginine	R	D-Arg, Lys, D-Lys, homo-Arg, D-homo-Arg, Met, Ile, D-Met, D-Ile, Orn, D-Orn
Asparagine	N	D-Asn, Asp, D-Asp, Glu, D-Glu, Gln, D-Gln
-Aspartic-Acid-	D	-D-Asp, D-Asn, Asn, Glu, D-Glu, Gln, D-Gln
Cysteine	С	D-Cys, S-Me-Cys, Met, D-Met, Thr, D-Thr
Glutamine	Q	D-Gln, Asn, D-Asn, Glu, D-Glu, Asp, D-Asp
Glutamic Acid	Е	D-Glu, D-Asp, Asp, Asn, D-Asn, Gln, D-Gln
Glycine	G	Ala, D-Ala, Pro, D-Pro, β-Ala, Acp
Isoleucine	I	D-Ile, Val, D-Val, Leu, D-Leu, Met, D-Met
Leucine	L	D-Leu, Val, D-Val, Leu, D-Leu, Met, D-Met
Lysine	K	D-Lys, Arg, D-Arg, homo-Arg, D-homo-Arg, Met, D-Met, Ile, D-Ile, Orn, D-Orn
Methionine	М	D-Met, S-Me-Cys, Ile, D-Ile, Leu, D-Leu, Val, D-Val
Phenylalanine	F	D-Phe, Tyr, D-Thr, L-Dopa, His, D-His, Trp, D-Trp, Trans-3,4, or 5-phenylproline, cis-3,4, or 5-phenylproline
Proline	P	D-Pro, L-I-thioazolidine-4-carboxylic acid, D-or L-1-oxazolidine-4-carboxylic acid
Serine	S	D-Ser, Thr, D-Thr, allo-Thr, Met, D-Met, Met(O), D-Met(O), L-Cys, D-Cys
Threonine	T	D-Thr, Ser, D-Ser, allo-Thr, Met, D-Met, Met(O), D-Met(O), Val, D-Val
Tyrosine	Y	D-Tyr, Phe, D-Phe, L-Dopa, His, D-His
Valine	V	D-Val, Leu, D-Leu, Ile, D-Ile, Met, D-Met

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Other analogs within the invention are those with modifications which increase peptide stability; such analogs may contain, for example, one or more non-peptide bonds (which replace the peptide bonds) in the peptide sequence. Also included are: analogs that include residues other than naturally occurring L-amino acids, e.g., D-amino acids or non-naturally occurring or synthetic amino acids, e.g., β or γ amino acids; and cyclic analogs.

As used herein, the term "fragment", as applied to an *H. pylori* analog, will ordinarily be at least about 20 residues, more typically at least about 40 residues, preferably at least about 60 residues in length. Fragments of *H. pylori* polypeptides can be generated by methods known to those skilled in the art. The ability of a candidate fragment to exhibit a biological activity of *H. pylori* polypeptide can be assessed by methods known to those skilled in the art as described herein. Also included are *H. pylori* polypeptides containing residues that are not required for biological activity of the peptide or that result from alternative mRNA splicing or alternative protein processing events.

An "immunogenic component" as used herein is a moiety, such as an *H. pylori* polypeptide, analog or fragment thereof, that is capable of eliciting a humoral and/or cellular immune response in a host animal alone or in combination with an adjuvant.

An "antigenic component" as used herein is a moiety, such as an *H. pylori* polypeptide, analog or fragment thereof, that is capable of binding to a specific antibody with sufficiently high affinity to form a detectable antigen-antibody complex.

As used herein, the term "transgene" means a nucleic acid (encoding, e.g., one or more polypeptides), which is partly or entirely heterologous, i.e., foreign, to the transgenic animal or cell into which it is introduced, or, is homologous to an endogenous gene of the transgenic animal or cell into which it is introduced, but which is designed to be inserted, or is inserted, into the cell's genome in such a way as to alter the genome of the cell into which it is inserted (e.g., it is inserted at a location which differs from that of the natural gene or its insertion results in a knockout). A transgene can include one or more transcriptional regulatory sequences and any other nucleic acid, such as introns, that may be necessary for optimal expression of the selected nucleic acid, all operably linked to the selected nucleic acid, and may include an enhancer sequence.

As used herein, the term "transgenic cell" refers to a cell containing a transgene.

As used herein, a "transgenic animal" is any animal in which one or more, and preferably essentially all, of the cells of the animal includes a transgene. The transgene can be introduced into the cell, directly or indirectly by introduction into a precursor of the cell, by way of deliberate genetic manipulation, such as by a process of transformation of competent cells or by microinjection or by infection with a

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recombinant virus. This molecule may be integrated within a chromosome, or it may be extrachromosomally replicating DNA.

The term "antibody" as used herein is intended to include fragments thereof which are specifically reactive with *H. pylori* polypeptides.

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As used herein, the term "cell-specific promoter" means a DNA sequence that serves as a promoter, i.e., regulates expression of a selected DNA sequence operably linked to the promoter, and which effects expression of the selected DNA sequence in specific cells of a tissue. The term also covers so-called "leaky" promoters, which regulate expression of a selected DNA primarily in one tissue, but cause expression in other tissues as well.

Misexpression, as used herein, refers to a non-wild type pattern of gene expression. It includes: expression at non-wild type levels, i.e., over or under expression; a pattern of expression that differs from wild type in terms of the time or stage at which the gene is expressed, e.g., increased or decreased expression (as compared with wild type) at a predetermined developmental period or stage; a pattern of expression that differs from wild type in terms of decreased expression (as compared with wild type) in a predetermined cell type or tissue type; a pattern of expression that differs from wild type in terms of the splicing size, amino acid sequence, post-transitional modification, or biological activity of the expressed polypeptide; a pattern of expression that differs from wild type in terms of the effect of an environmental stimulus or extracellular stimulus on expression of the gene, e.g., a pattern of increased or decreased expression (as compared with wild type) in the presence of an increase or decrease in the strength of the stimulus.

As used herein, "host cells" and other such terms denoting microorganisms or higher eukaryotic cell lines cultured as unicellular entities refers to cells which can become or have been used as recipients for a recombinant vector or other transfer DNA, and include the progeny of the original cell which has been transfected. It is understood by individuals skilled in the art that the progeny of a single parental cell may not necessarily be completely identical in genomic or total DNA compliment to the original parent, due to accident or deliberate mutation.

As used herein, the term "control sequence" refers to a nucleic acid having a base sequence which is recognized by the host organism to effect the expression of encoded sequences to which they are ligated. The nature of such control sequences differs depending upon the host organism; in prokaryotes, such control sequences generally include a promoter, ribosomal binding site, terminators, and in some cases operators; in eukaryotes, generally such control sequences include promoters, terminators and in some instances, enhancers. The term control sequence is intended to include at a

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minimum, all components whose presence is necessary for expression, and may also include additional components whose presence is advantageous, for example, leader sequences.

As used herein, the term "operably linked" refers to sequences joined or ligated to function in their intended manner. For example, a control sequence is operably linked to coding sequence by ligation in such a way that expression of the coding sequence is achieved under conditions compatible with the control sequence and host cell.

The metabolism of a substance, as used herein, means any aspect of the, expression, function, action, or regulation of the substance. The metabolism of a substance includes modifications, e.g., covalent or non-covalent modifications of the substance. The metabolism of a substance includes modifications, e.g., covalent or non-covalent modification, the substance induces in other substances. The metabolism of a substance also includes changes in the distribution of the substance. The metabolism of a substance includes changes the substance induces in the distribution of other substances.

A "sample" as used herein refers to a biological sample, such as, for example, tissue or fluid isloated from an individual (including without limitation plasma, serum, cerebrospinal fluid, lymph, tears, saliva and tissue sections) or from *in vitro* cell culture constituents, as well as samples from the environment.

The practice of the invention will employ, unless otherwise indicated, conventional techniques of chemistry, molecular biology, microbiology, recombinant DNA, and immunology, which are within the skill of the art. Such techniques are explained fully in the literature. See e.g., Sambrook, Fritsch, and Maniatis, Molecular Cloning: Laboratory Manual 2nd ed. (1989); DNA Cloning, Volumes I and II (D.N Glover ed. 1985); Oligonucleotide Synthesis (M.J. Gait ed, 1984); Nucleic Acid Hybridization (B.D. Hames & S.J. Higgins eds. 1984); the series, Methods in Enzymology (Academic Press, Inc.), particularly Vol. 154 and Vol. 155 (Wu and Grossman, eds.) and PCR-A Practical Approach (McPherson, Quirke, and Taylor, eds., 1991).

I. Isolation of Nucleic Acids of H. pylori and Uses Therefor

H. pylori Genomic Sequence

This invention provides nucleotide sequences of the genome of *H. pylori* which thus comprises a DNA sequence library of *H. pylori* genomic DNA. The detailed description that follows provides nucleotide sequences of *H. pylori*, and also describes how the sequences were obtained and how ORFs and protein-coding sequences were

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identified. Also described are methods of using the disclosed *H. pylori* sequences in methods including diagnostic and therapeutic applications. Furthermore, the library can be used as a database for identification and comparison of medically important sequences in this and other strains of *H. pylori*.

To determine the genomic sequence of *H. pylori*, DNA was isolated from a strain of *H. pylori* (ATCC # 55679; deposited by Genome Therapeutics Corporation, 100 Beaver Street, Waltham, MA 02154) and mechanically sheared by nebulization to a median size of 2 kb. Following size fractionation by gel electrophoresis, the fragments were blunt-ended, ligated to adapter oligonucleotides, and cloned into each of 20 different pMPX vectors (Rice et al., abstracts of Meeting of Genome Mapping and Sequencing, Cold Spring Harbor, NY, 5/11-5/15, 1994, p. 225) to construct a series of "shotgun" subclone libraries.

DNA sequencing was achieved using multiplex sequencing procedures essentially as disclosed in Church et al., 1988, *Science* 240:185; U.S. Patents No. 4,942,124 and 5,149,625). DNA was extracted from pooled cultures and subjected to chemical or enzymatic sequencing. Sequencing reactions were resolved by electrophoresis, and the products were transferred and covalently bound to nylon membranes. Finally, the membranes were sequentially hybridized with a series of labelled oligonucleotides complimentary to "tag" sequences present in the different shotgun cloning vectors. In this manner, a large number of sequences could be obtained from a single set of sequencing reactions. The cloning and sequencing procedures are described in more detail in the Exemplification.

Individual sequence reads obtained in this manner were assembled using the FALCONTM program (Church *et al.*, 1994, *Automated DNA Sequencing and Analysis*, J.C. Venter, ed., Academic Press) and PHRAP (P. Green, Abstracts of DOE Human Genome Program Contractor-Grantee Workshop V, Jan. 1996, p.157). The average contig length was about 3-4 kb.

A variety of approaches are used to order the contigs so as to obtain a continuous sequence representing the entire *H. pylori* genome. Synthetic oligonucleotides are designed that are complementary to sequences at the end of each contig. These oligonucleotides may be hybridized to libaries of *H. pylori* genomic DNA in, for example, lambda phage vectors or plasmid vectors to identify clones that contain sequences corresponding to the junctional regions between individual contigs. Such clones are then used to isolate template DNA and the same oligonucleotides are used as primers in polymerase chain reaction (PCR) to amplify junctional fragments, the nucleotide sequence of which is then determined.

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The *H. pylori* sequences were analyzed for the presence of open reading frames (ORFs) comprising at least 180 nucleotides. As a result of the analysis of ORFs based on stop-to-stop codon reads, it should be understood that these ORFs may not correspond to the ORF of a naturally-occurring *H. pylori* polypeptide. These ORFs may contain start codons which indicate the initiation of protein synthesis of a naturally-occurring *H. pylori* polypeptide. Such start codons within the ORFs provided herein can be identified by those of ordinary skill in the relevant art, and the resulting ORF and the encoded *H. pylori* polypeptide is within the scope of this invention. For example, within the ORFs a codon such as AUG or GUG (encoding methionine or valine) which is part of the initiation signal for protein synthesis can be identified and the ORF modified to correspond to a naturally-occurring *H. pylori* polypeptide. The predicted coding regions were defined by evaluating the coding potential of such sequences with the program GENEMARKTM (Borodovsky and McIninch, 1993, *Comp. Chem.* 17:123).

Other H. pylori Nucleic Acids

The nucleic acids of this invention may be obtained directly from the DNA of the above referenced *H. pylori* strain by using the polymerase chain reaction (PCR). See "PCR, A Practical Approach" (McPherson, Quirke, and Taylor, eds., IRL Press, Oxford, UK, 1991) for details about the PCR. High fidelity PCR can be used to ensure a faithful DNA copy prior to expression. In addition, the authenticity of amplified products can be checked by conventional sequencing methods. Clones carrying the desired sequences described in this invention may also be obtained by screening the libraries by means of the PCR or by hybridization of synthetic oligonucleotide probes to filter lifts of the library colonies or plaques as known in the art (see, e.g., Sambrook et al., Molecular Cloning, A Laboratory Manual 2nd edition, 1989, Cold Spring Harbor Press, NY).

It is also possible to obtain nucleic acids encoding *H. pylori* polypeptides from a cDNA library in accordance with protocols herein described. A cDNA encoding an *H. pylori* polypeptide can be obtained by isolating total mRNA from an appropriate strain. Double stranded cDNAs can then be prepared from the total mRNA. Subsequently, the cDNAs can be inserted into a suitable plasmid or viral (e.g., bacteriophage) vector using any one of a number of known techniques. Genes encoding *H. pylori* polypeptides can also be cloned using established polymerase chain reaction techniques in accordance with the nucleotide sequence information provided by the invention. The nucleic acids of the invention can be DNA or RNA. Preferred nucleic acids of the invention are contained in the Sequence Listing.

The nucleic acids of the invention can also be chemically synthesized using standard techniques. Various methods of chemically synthesizing polydeoxynucleotides

are known, including solid-phase synthesis which, like peptide synthesis, has been fully automated in commercially available DNA synthesizers (See e.g., Itakura et al. U.S. Patent No. 4,598,049; Caruthers et al. U.S. Patent No. 4,458,066; and Itakura U.S. Patent Nos. 4,401,796 and 4,373,071, incorporated by reference herein).

Nucleic acids isolated or synthesized in accordance with features of the present invention are useful, by way of example, without limitation, as probes, primers, capture ligands, antisense genes and for developing expression systems for the synthesis of proteins and peptides corresponding to such sequences. As probes, primers, capture ligands and antisense agents, the nucleic acid normally consists of all or part (approximately twenty or more nucleotides for specificity as well as the ability to form stable hybridization products) of the nucleic acids of the invention contained in the Sequence Listing. These uses are described in further detail below.

Probes

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A nucleic acid isolated or synthesized in accordance with the sequence of the invention contained in the Sequence Listing can be used as a probe to specifically detect *H. pylori*. With the sequence information set forth in the present application, sequences of twenty or more nucleotides are identified which provide the desired inclusivity and exclusivity with respect to *H. pylori*, and extraneous nucleic acids likely to be encountered during hybridization conditions. More preferably, the sequence will comprise at least twenty to thirty nucleotides to convey stability to the hybridization product formed between the probe and the intended target molecules.

Sequences larger than 1000 nucleotides in length are difficult to synthesize but can be generated by recombinant DNA techniques. Individuals skilled in the art will readily recognize that the nucleic acids, for use as probes, can be provided with a label to facilitate detection of a hybridization product.

Nucleic acid isolated and synthesized in accordance with the sequence of the invention contained in the Sequence Listing can also be useful as probes to detect homologous regions (especially homologous genes) of other *Helicobacter* species using appropriate stringency hybridization conditions as described herein.

Capture Ligand

For use as a capture ligand, the nucleic acid selected in the manner described above with respect to probes, can be readily associated with a support. The manner in which nucleic acid is associated with supports is well known. Nucleic acid having twenty or more nucleotides in a sequence of the invention contained in the Sequence Listing have utility to separate *H. pylori* nucleic acid from the nucleic acid of each other and other organisms. Nucleic acid having twenty or more nucleotides in a sequence of the invention contained in the Sequence Listing can also have utility to separate other

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Helicobacter species from each other and from other organisms. Preferably, the sequence will comprise at least twenty nucleotides to convey stability to the hybridization product formed between the probe and the intended target molecules. Sequences larger than 1000 nucleotides in length are difficult to synthesize but can be generated by recombinant DNA techniques.

Primers

Nucleic acid isolated or synthesized in accordance with the sequences described herein have utility as primers for the amplification of H. pylori nucleic acid. These nucleic acids may also have utility as primers for the amplification of nucleic acids in other Helicobacter species. With respect to polymerase chain reaction (PCR) techniques, nucleic acid sequences of $\geq 10\text{-}15$ nucleotides of the invention contained in the Sequence Listing have utility in conjunction with suitable enzymes and reagents to create copies of H. pylori nucleic acid. More preferably, the sequence will comprise twenty or more nucleotides to convey stability to the hybridization product formed between the primer and the intended target molecules. Binding conditions of primers greater than 100 nucleotides are more difficult to control to obtain specificity. High fidelity PCR can be used to ensure a faithful DNA copy prior to expression. In addition, amplified products can be checked by conventional sequencing methods.

The copies can be used in diagnostic assays to detect specific sequences, including genes from *H. pylori* and/or other *Helicobacter* species. The copies can also be incorporated into cloning and expression vectors to generate polypeptides corresponding to the nucleic acid synthesized by PCR, as is described in greater detail herein.

<u>Antisense</u>

Nucleic acid or nucleic acid-hybridizing derivatives isolated or synthesized in accordance with the sequences described herein have utility as antisense agents to prevent the expression of *H. pylori* genes. These sequences also have utility as antisense agents to prevent expression of genes of other *Helicobacter* species.

In one embodiment, nucleic acid or derivatives corresponding to *H. pylori* nucleic acids is loaded into a suitable carrier such as a liposome or bacteriophage for introduction into bacterial cells. For example, a nucleic acid having twenty or more nucleotides is capable of binding to bacteria nucleic acid or bacteria messenger RNA. Preferably, the antisense nucleic acid is comprised of 20 or more nucleotides to provide necessary stability of a hybridization product of non-naturally occurring nucleic acid and bacterial nucleic acid and/or bacterial messenger RNA. Nucleic acid having a sequence greater than 1000 nucleotides in length is difficult to synthesize but can be generated by recombinant DNA techniques. Methods for loading antisense nucleic acid in liposomes

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is known in the art as exemplified by U.S. Patent 4,241,046 issued December 23, 1980 to Papahadjopoulos et al.

II. Expression of H. pylori Nucleic Acids

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Nucleic acid isolated or synthesized in accordance with the sequences described herein have utility to generate polypeptides. The nucleic acid of the invention exemplified in the Sequence Listing or fragments of said nucleic acid encoding active portions of *H. pylori* polypeptides can be cloned into suitable vectors or used to isolate nucleic acid. The isolated nucleic acid is combined with suitable DNA linkers and cloned into a suitable vector.

The function of a specific gene or operon can be ascertained by expression in a bacterial strain under conditions where the activity of the gene product(s) specified by the gene or operon in question can be specifically measured. Alternatively, a gene product may be produced in large quantities in an expressing strain for use as an antigen, an industrial reagent, for structural studies, etc. This expression can be accomplished in a mutant strain which lacks the activity of the gene to be tested, or in a strain that does not produce the same gene product(s). This includes, but is not limited to other Helicobacter strains, or other bacterial strains such as E. coli, Norcardia, Corynebacterium, Campylobacter, and Streptomyces species. In some cases the expression host will utilize the natural Helicobacter promoter whereas in others, it will be necessary to drive the gene with a promoter sequence derived from the expressing organism (e.g., an E. coli beta-galactosidase promoter for expression in E. coli).

To express a gene product using the natural *H. pylori* promoter, a procedure such as the following can be used. A restriction fragment containing the gene of interest, together with its associated natural promoter element and regulatory sequences (identified using the DNA sequence data) is cloned into an appropriate recombinant plasmid containing an origin of replication that functions in the host organism and an appropriate selectable marker. This can be accomplished by a number of procedures known to those skilled in the art. It is most preferably done by cutting the plasmid and the fragment to be cloned with the same restriction enzyme to produce compatible ends that can be ligated to join the two pieces together. The recombinant plasmid is introduced into the host organism by, for example, electroporation and cells containing the recombinant plasmid are identified by selection for the marker on the plasmid. Expression of the desired gene product is detected using an assay specific for that gene product.

In the case of a gene that requires a different promoter, the body of the gene (coding sequence) is specifically excised and cloned into an appropriate expression

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plasmid. This subcloning can be done by several methods, but is most easily accomplished by PCR amplification of a specific fragment and ligation into an expression plasmid after treating the PCR product with a restriction enzyme or exonuclease to create suitable ends for cloning.

A suitable host cell for expression of a gene can be any procaryotic or eucaryotic cell. For example, an *H. pylori* polypeptide can be expressed in bacterial cells such as *E. coli*, insect cells (baculovirus), yeast, or mammalian cells such as Chinese hamster ovary cell (CHO). Other suitable host cells are known to those skilled in the art.

Expression in eucaryotic cells such as mammalian, yeast, or insect cells can lead to partial or complete glycosylation and/or formation of relevant inter- or intra-chain disulfide bonds of a recombinant peptide product. Examples of vectors for expression in yeast S. cerivisae include pYepSec1 (Baldari. et al., (1987) Embo J. 6:229-234), pMFa (Kurjan and Herskowitz, (1982) Cell 30:933-943), pJRY88 (Schultz et al., (1987) Gene 54:113-123), and pYES2 (Invitrogen Corporation, San Diego, CA). Baculovirus vectors available for expression of proteins in cultured insect cells (SF 9 cells) include the pAc series (Smith et al., (1983) Mol. Cell Biol. 3:2156-2165) and the pVL series (Lucklow, V.A., and Summers, M.D., (1989) Virology 170:31-39). Generally, COS cells (Gluzman, Y., (1981) Cell 23:175-182) are used in conjunction with such vectors as pCDM 8 (Aruffo, A. and Seed, B., (1987) Proc. Natl. Acad. Sci. USA 84:8573-8577) for transient amplification/expression in mammalian cells, while CHO (dhfr- Chinese Hamster Ovary) cells are used with vectors such as pMT2PC (Kaufman et al. (1987), EMBO J. 6:187-195) for stable amplification/expression in mammalian cells. Vector DNA can be introduced into mammalian cells via conventional techniques such as calcium phosphate or calcium chloride co-precipitation, DEAE-dextran-mediated transfection, or electroporation. Suitable methods for transforming host cells can be found in Sambrook et al. (Molecular Cloning: A Laboratory Manual, 2nd Edition, Cold Spring Harbor Laboratory press (1989)), and other laboratory textbooks.

Expression in procaryotes is most often carried out in *E. coli* with either fusion or non-fusion inducible expression vectors. Fusion vectors usually add a number of NH₂ terminal amino acids to the expressed target gene. These NH₂ terminal amino acids often are referred to as a reporter group. Such reporter groups usually serve two purposes: 1) to increase the solubility of the target recombinant protein; and 2) to aid in the purification of the target recombinant protein by acting as a ligand in affinity purification. Often, in fusion expression vectors, a proteolytic cleavage site is introduced at the junction of the reporter group and the target recombinant protein to enable separation of the target recombinant protein from the reporter group subsequent to purification of the fusion protein. Such enzymes, and their cognate recognition

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sequences, include Factor Xa, thrombin and enterokinase. Typical fusion expression vectors include pGEX (Amrad Corp., Melbourne, Australia), pMAL (New England Biolabs, Beverly, MA) and pRIT5 (Pharmacia, Piscataway, NJ) which fuse glutathione S-transferase, maltose E binding protein, or protein A, respectively, to the target recombinant protein. A preferred reporter group is poly(His), which may be fused to the amino or carboxy terminus of the protein and which renders the recombinant fusion protein easily purifiable by metal chelate chromatography.

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Inducible non-fusion expression vectors include pTrc (Amann et al., (1988) Gene 69:301-315) and pET11d (Studier et al., Gene Expression Technology: Methods in Enzymology 185, Academic Press, San Diego, California (1990) 60-89). While target gene expression relies on host RNA polymerase transcription from the hybrid trp-lac fusion promoter in pTrc, expression of target genes inserted into pET11d relies on transcription from the T7 gn10-lac 0 fusion promoter mediated by coexpressed viral RNA polymerase (T7 gn1). This viral polymerase is supplied by host strains BL21(DE3) or HMS174(DE3) from a resident λ prophage harboring a T7 gn1 under the transcriptional control of the lacUV 5 promoter.

For example, a host cell transfected with a nucleic acid vector directing expression of a nucleotide sequence encoding an *H. pylori* polypeptide can be cultured under appropriate conditions to allow expression of the polypeptide to occur. The polypeptide may be secreted and isolated from a mixture of cells and medium containing the peptide. Alternatively, the polypeptide may be retained cytoplasmically and the cells harvested, lysed and the protein isolated. A cell culture includes host cells, media and other byproducts. Suitable media for cell culture are well known in the art.

Polypeptides of the invention can be isolated from cell culture medium, host cells, or both using techniques known in the art for purifying proteins including ion-exchange chromatography, gel filtration chromatography, ultrafiltration, electrophoresis, and immunoaffinity purification with antibodies specific for such polypeptides.

Additionally, in many situations, polypeptides can be produced by chemical cleavage of a native protein (e.g., tryptic digestion) and the cleavage products can then be purified by standard techniques.

In the case of membrane bound proteins, these can be isolated from a host cell by contacting a membrane-associated protein fraction with a detergent forming a solubilized complex, where the membrane-associated protein is no longer entirely embedded in the membrane fraction and is solubilized at least to an extent which allows it to be chromatographically isolated from the membrane fraction. Several different criteria are used for choosing a detergent suitable for solubilizing these complexes. For example, one property considered is the ability of the detergent to solubilize the H.

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pylori protein within the membrane fraction at minimal denaturation of the membraneassociated protein allowing for the activity or functionality of the membrane-associated protein to return upon reconstitution of the protein. Another property considered when selecting the detergent is the critical micelle concentration (CMC) of the detergent in that the detergent of choice preferably has a high CMC value allowing for ease of removal after reconstitution. A third property considered when selecting a detergent is the hydrophobicity of the detergent. Typically, membrane-associated proteins are very hydrophobic and therefore detergents which are also hydrophobic, e.g., the triton series, would be useful for solubilizing the hydrophobic proteins. Another property important to a detergent can be the capability of the detergent to remove the H. pylori protein with minimal protein-protein interaction facilitating further purification. A fifth property of the detergent which should be considered is the charge of the detergent. For example, if it is desired to use ion exchange resins in the purification process then preferably detergent should be an uncharged detergent. Chromatographic techniques which can be used in the final purification step are known in the art and include hydrophobic interaction, lectin affinity, ion exchange, dye affinity and immunoaffinity.

One strategy to maximize recombinant *H. pylori* peptide expression in *E. coli* is to express the protein in a host bacteria with an impaired capacity to proteolytically cleave the recombinant protein (Gottesman, S., Gene Expression Technology: Methods in Enzymology 185, Academic Press, San Diego, California (1990) 119-128). Another strategy would be to alter the nucleic acid encoding an *H. pylori* peptide to be inserted into an expression vector so that the individual codons for each amino acid would be those preferentially utilized in highly expressed *E. coli* proteins (Wada et al., (1992) *Nuc. Acids Res.* 20:2111-2118). Such alteration of nucleic acids of the invention can be carried out by standard DNA synthesis techniques.

The nucleic acids of the invention can also be chemically synthesized using standard techniques. Various methods of chemically synthesizing polydeoxynucleotides are known, including solid-phase synthesis which, like peptide synthesis, has been fully automated in commercially available DNA synthesizers (See, e.g., Itakura et al. U.S. Patent No. 4,598,049; Caruthers et al. U.S. Patent No. 4,458,066; and Itakura U.S. Patent Nos. 4,401,796 and 4,373,071, incorporated by reference herein).

III. H. pylori Polypeptides

This invention encompasses isolated *H. pylori* polypeptides encoded by the disclosed *H. pylori* genomic sequences, including the polypeptides of the invention contained in the Sequence Listing. Polypeptides of the invention are preferably at least 5 amino acid residues in length. Using the DNA sequence information provided herein,

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the amino acid sequences of the polypeptides encompassed by the invention can be deduced using methods well-known in the art. It will be understood that the sequence of an entire nucleic acid encoding an *H. pylori* polypeptide can be isolated and identified based on an ORF that encodes only a fragment of the cognate protein-coding region. This can be acheived, for example, by using the isolated nucleic acid encoding the ORF, or fragments thereof, to prime a polymerase chain reaction with genomic *H. pylori* DNA as template; this is followed by sequencing the amplified product.

The polypeptides of the invention can be isolated from wild-type or mutant *H. pylori* cells or from heterologous organisms or cells (including, but not limited to, bacteria, yeast, insect, plant and mammalian cells) into which an *H. pylori* nucleic acid has been introduced and expressed. In addition, the polypeptides can be part of recombinant fusion proteins.

H. pylori polypeptides of the invention can be chemically synthesized using commercially automated procedures such as those referenced herein.

IV. Identification of Nucleic Acids Encoding Vaccine Components and Targets for Agents Effective Against H. pylori

The disclosed *H. pylori* genome sequence includes segments that direct the synthesis of ribonucleic acids and polypeptides, as well as origins of replication, promoters, other types of regulatory sequences, and intergenic nucleic acids. The invention encompasses nucleic acids encoding immunogenic components of vaccines and targets for agents effective against *H. pylori*. Identification of said immunogenic components involved in the determination of the function of the disclosed sequences can be achieved using a variety of approaches. Non-limiting examples of these approaches are described briefly below.

Homology to known sequences: Computer-assisted comparison of the disclosed *H. pylori* sequences with previously reported sequences present in publicly available databases is useful for identifying functional *H. pylori* nucleic acid and polypeptide sequences. It will be understood that protein-coding sequences, for example, may be compared as a whole, and that a high degree of sequence homology between two proteins (such as, for example, >80-90%) at the amino acid level indicates that the two proteins also possess some degree of functional homology, such as, for example, among enzymes involved in metabolism, DNA synthesis, or cell wall synthesis, and proteins involved in transport, cell division, etc. In addition, many structural features of particular protein classes have been identified and correlate with specific consensus sequences, such as, for example, binding domains for nucleotides, DNA, metal ions, and other small molecules; sites for covalent modifications such as phosphorylation,

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acylation, and the like; sites of protein:protein interactions, etc. These consensus sequences may be quite short and thus may represent only a fraction of the entire protein-coding sequence. Identification of such a feature in an *H. pylori* sequence is therefore useful in determining the function of the encoded protein and identifying useful targets of antibacterial drugs.

Of particular relevance to the present invention are structural features that are common to secretory, transmembrane, and surface proteins, including secretion signal peptides and hydrophobic transmembrane domains. *H. pylori* proteins identified as containing putative signal sequences and/or transmembrane domains are useful as immunogenic components of vaccines.

<u>Identification of essential genes:</u> Nucleic acids that encode proteins essential for growth or viability of *H. pylori* are preferred drug targets. *H. pylori* genes can be tested for their biological relevance to the organism by examining the effect of deleting and/or disrupting the genes, i.e., by so-called gene "knockout", using techniques known to those skilled in the relevant art. In this manner, essential genes may be identified.

Strain-specific sequences: Because of the evolutionary relationship between different *H. pylori* strains, it is believed that the presently disclosed *H. pylori* sequences are useful for identifying, and/or discriminating between, previously known and new *H. pylori* strains. It is believed that other *H. pylori* strains will exhibit at least 70% sequence homology with the presently disclosed sequence. Systematic and routine analyses of DNA sequences derived from samples containing *H. pylori* strains, and comparison with the present sequence allows for the identification of sequences that can be used to discriminate between strains, as well as those that are common to all *H. pylori* strains. In one embodiment, the invention provides nucleic acids, including probes, and peptide and polypeptide sequences that discriminate between different strains of *H. pylori*. Strain-specific components can also be identified functionally by their ability to elicit or react with antibodies that selectively recognize one or more *H. pylori* strains.

In another embodiment, the invention provides nucleic acids, including probes, and peptide and polypeptide sequences that are common to all *H. pylori* strains but are *not* found in other bacterial species.

Specific Example: Determination Of Candidate Protein Antigens For Antibody And Vaccine Development

The selection of candidate protein antigens for vaccine development can be derived from the nucleic acids encoding *H. pylori* polypeptides. First, the ORF's can be analyzed for homology to other known exported or membrane proteins and analyzed using the discriminant analysis described by Klein, et al. (Klein, P., Kanehsia, M., and

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DeLisi, C. (1985) Biochimica et Biophysica Acta 815, 468-476) for predicting exported and membrane proteins.

Homology searches can be performed using the BLAST algorithm contained in the Wisconsin Sequence Analysis Package (Genetics Computer Group, University Research Park, 575 Science Drive, Madison, WI 53711) to compare each predicted ORF amino acid sequence with all sequences found in the current GenBank, SWISS-PROT and PIR databases. BLAST searches for local alignments between the ORF and the databank sequences and reports a probability score which indicates the probability of finding this sequence by chance in the database. ORF's with significant homology (e.g. probabilities lower than 1x10-6 that the homology is only due to random chance) to membrane or exported proteins represent protein antigens for vaccine development. Possible functions can be provided to *H. pylori* genes based on sequence homology to genes cloned in other organisms.

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Discriminant analysis (Klein, et al. supra) can be used to examine the ORF amino acid sequences. This algorithm uses the intrinsic information contained in the ORF amino acid sequence and compares it to information derived from the properties of known membrane and exported proteins. This comparison predicts which proteins will be exported, membrane associated or cytoplasmic. ORF amino acid sequences identified as exported or membrane associated by this algorithm are likely protein antigens for vaccine development.

Surface exposed outer membrane proteins are likely to represent the best antigens to provide a protective immune response against *H. pylori*. Among the algorithms that can be used to aid in prediction of these outer membrane proteins include the presence of an amphipathic beta-sheet region at their C-terminus. This region which has been detected in a large number of outer membrane proteins in Gram negative bacteria is often characterized by hydrophobic residues (Phe or Tyr) clustered at alternating positions from the C-terminus (e.g., see Figure 5, block F; Figure 7, block E). Importantly, these sequences have not been detected at the C-termini of periplasmic proteins, thus allowing preliminary distinction between these classes of proteins based on primary sequence data. This phenomenon has been reported previously by Struyve et al. (*J. Mol. Biol.* 218:141-148, 1991).

Also illustrated in Figure 5 are additional amino acid sequence motifs found in many outer membrane proteins of *H. pylori*. The amino acid sequence alignment in Figure 5 depicts portions of the sequence of five *H. pylori* proteins (depicted in the single letter amino acid code) labeled with their amino acid Sequence ID Numbers and shown N-terminal to C-terminal, left to right. Five or six distinct blocks (labeled A through E or F) of similar amino acid residues are found including the distinctive

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hydrophobic residues (Phe or Tyr; F or Y according to the single letter code for amino acid residues) frequently found at positions near the C-terminus of outer membrane proteins. The presence of several shared motifs clearly establishes the similarity between members of this group of proteins.

Additional amino acid alignments for four outer membrane proteins isolated from *H. pylori* are depicted in Figure 6.

Outer membrane proteins isolated from *H. pylori* frequently share additional motifs as depicted for two proteins in Figure 7 which also share the C-terminal hydrophobic residues, and as depicted for two proteins in Figure 8 which do not share the C-terminal hydrophobic residue motif but share a different C-terminal motif.

One skilled in the art would know that these shared sequence motifs are highly significant and establish a similarity among this group of proteins.

Infrequently it is not possible to distinguish between multiple possible nucleotides at a given position in the nucleic acid sequence. In those cases the ambiguities are denoted by an extended alphabet as follows:

These are the official IUPAC-IUB single-letter base codes

Code	Base Description	
G	Guanine	
Α	Adenine	
T	Thymine	
С	Cytosine	
R	Purine	(A or G)
Y	Pyrimidine	(C or T or U)
M	Amino	(A or C)
K	Ketone	(G or T)
S	Strong interaction	(C or G)
W	Weak interaction	(A or T)
Н	Not-G	(A or C or T)
В	Not-A	(C or G or T)
V	Not-T (not-U)	(A or C or G)
D	Not-C	(A or G or T)
N	Any	(A or C or G or T)

The amino acid translations of this invention account for the ambiguity in the nucleic acid sequence by translating the ambiguous codon as the letter "X". In all cases,

the permissible amino acid residues at a position are clear from an examination of the nucleic acid sequence based on the standard genetic code.

V. Production of Fragments and Analogs of H. pylori Nucleic Acids and Polypeptides

Based on the discovery of the *H. pylori* gene products of the invention provided in the Sequence Lsiting, one skilled in the art can alter the disclosed structure (of *H. pylori* genes), e.g., by producing fragments or analogs, and test the newly produced structures for activity. Examples of techniques known to those skilled in the relevant art which allow the production and testing of fragments and analogs are discussed below. These, or analogous methods can be used to make and screen libraries of polypeptides, e.g., libraries of random peptides or libraries of fragments or analogs of cellular proteins for the ability to bind *H. pylori* polypeptides. Such screens are useful for the identification of inhibitors of *H. pylori*.

Generation of Fragments

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Fragments of a protein can be produced in several ways, e.g., recombinantly, by proteolytic digestion, or by chemical synthesis. Internal or terminal fragments of a polypeptide can be generated by removing one or more nucleotides from one end (for a terminal fragment) or both ends (for an internal fragment) of a nucleic acid which encodes the polypeptide. Expression of the mutagenized DNA produces polypeptide fragments. Digestion with "end-nibbling" endonucleases can thus generate DNA's which encode an array of fragments. DNA's which encode fragments of a protein can also be generated by random shearing, restriction digestion or a combination of the above-discussed methods.

Fragments can also be chemically synthesized using techniques known in the art such as conventional Merrifield solid phase f-Moc or t-Boc chemistry. For example, peptides of the present invention may be arbitrarily divided into fragments of desired length with no overlap of the fragments, or divided into overlapping fragments of a desired length.

Alteration of Nucleic Acids and Polypeptides: Random Methods

Amino acid sequence variants of a protein can be prepared by random mutagenesis of DNA which encodes a protein or a particular domain or region of a protein. Useful methods include PCR mutagenesis and saturation mutagenesis. A library of random amino acid sequence variants can also be generated by the synthesis of a set of degenerate oligonucleotide sequences. (Methods for screening proteins in a library of variants are elsewhere herein).

(A) PCR Mutagenesis

In PCR mutagenesis, reduced Taq polymerase fidelity is used to introduce random mutations into a cloned fragment of DNA (Leung et al., 1989, *Technique* 1:11-15). The DNA region to be mutagenized is amplified using the polymerase chain reaction (PCR) under conditions that reduce the fidelity of DNA synthesis by Taq DNA polymerase, e.g., by using a dGTP/dATP ratio of five and adding Mn²⁺ to the PCR reaction. The pool of amplified DNA fragments are inserted into appropriate cloning vectors to provide random mutant libraries.

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(B) Saturation Mutagenesis

Saturation mutagenesis allows for the rapid introduction of a large number of single base substitutions into cloned DNA fragments (Mayers et al., 1985, Science 229:242). This technique includes generation of mutations, e.g., by chemical treatment or irradiation of single-stranded DNA in vitro, and synthesis of a complimentary DNA strand. The mutation frequency can be modulated by modulating the severity of the treatment, and essentially all possible base substitutions can be obtained. Because this procedure does not involve a genetic selection for mutant fragments both neutral substitutions, as well as those that alter function, are obtained. The distribution of point mutations is not biased toward conserved sequence elements.

(C) Degenerate Oligonucleotides

A library of homologs can also be generated from a set of degenerate oligonucleotide sequences. Chemical synthesis of a degenerate sequences can be carried out in an automatic DNA synthesizer, and the synthetic genes then ligated into an appropriate expression vector. The synthesis of degenerate oligonucleotides is known in the art (see for example, Narang, SA (1983) Tetrahedron 39:3; Itakura et al. (1981) Recombinant DNA, Proc 3rd Cleveland Sympos. Macromolecules, ed. AG Walton, Amsterdam: Elsevier pp273-289; Itakura et al. (1984) Annu. Rev. Biochem. 53:323; Itakura et al. (1984) Science 198:1056; Ike et al. (1983) Nucleic Acid Res. 11:477. Such techniques have been employed in the directed evolution of other proteins (see, for example, Scott et al. (1990) Science 249:386-390; Roberts et al. (1992) PNAS 89:2429-2433; Devlin et al. (1990) Science 249: 404-406; Cwirla et al. (1990) PNAS 87: 6378-6382; as well as U.S. Patents Nos. 5,223,409, 5,198,346, and 5,096,815).

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Alteration of Nucleic Acids and Polypeptides: Methods for Directed Mutagenesis

Non-random or directed, mutagenesis techniques can be used to provide specific sequences or mutations in specific regions. These techniques can be used to create variants which include, e.g., deletions, insertions, or substitutions, of residues of the known amino acid sequence of a protein. The sites for mutation can be modified individually or in series, e.g., by (1) substituting first with conserved amino acids and then with more radical choices depending upon results achieved, (2) deleting the target residue, or (3) inserting residues of the same or a different class adjacent to the located site, or combinations of options 1-3.

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(A) Alanine Scanning Mutagenesis

Alanine scanning mutagenesis is a useful method for identification of certain residues or regions of the desired protein that are preferred locations or domains for mutagenesis, Cunningham and Wells (*Science* 244:1081-1085, 1989). In alanine scanning, a residue or group of target residues are identified (e.g., charged residues such as Arg, Asp, His, Lys, and Glu) and replaced by a neutral or negatively charged amino acid (most preferably alanine or polyalanine). Replacement of an amino acid can affect the interaction of the amino acids with the surrounding aqueous environment in or outside the cell. Those domains demonstrating functional sensitivity to the substitutions are then refined by introducing further or other variants at or for the sites of substitution. Thus, while the site for introducing an amino acid sequence variation is predetermined, the nature of the mutation per se need not be predetermined. For example, to optimize the performance of a mutation at a given site, alanine scanning or random mutagenesis may be conducted at the target codon or region and the expressed desired protein subunit variants are screened for the optimal combination of desired activity.

(B) Oligonucleotide-Mediated Mutagenesis

Oligonucleotide-mediated mutagenesis is a useful method for preparing substitution, deletion, and insertion variants of DNA, see, e.g., Adelman et al., (DNA 2:183, 1983). Briefly, the desired DNA is altered by hybridizing an oligonucleotide encoding a mutation to a DNA template, where the template is the single-stranded form of a plasmid or bacteriophage containing the unaltered or native DNA sequence of the desired protein. After hybridization, a DNA polymerase is used to synthesize an entire second complementary strand of the template that will thus incorporate the oligonucleotide primer, and will code for the selected alteration in the desired protein DNA. Generally, oligonucleotides of at least 25 nucleotides in length are used. An optimal oligonucleotide will have 12 to 15 nucleotides that are completely

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complementary to the template on either side of the nucleotide(s) coding for the mutation. This ensures that the oligonucleotide will hybridize properly to the single-stranded DNA template molecule. The oligonucleotides are readily synthesized using techniques known in the art such as that described by Crea et al. (*Proc. Natl. Acad. Sci.* USA, 75: 5765[1978]).

(C) Cassette Mutagenesis

Another method for preparing variants, cassette mutagenesis, is based on the technique described by Wells et al. (Gene, 34:315[1985]). The starting material is a plasmid (or other vector) which includes the protein subunit DNA to be mutated. The codon(s) in the protein subunit DNA to be mutated are identified. There must be a unique restriction endonuclease site on each side of the identified mutation site(s). If no such restriction sites exist, they may be generated using the above-described oligonucleotide-mediated mutagenesis method to introduce them at appropriate locations in the desired protein subunit DNA. After the restriction sites have been introduced into the plasmid, the plasmid is cut at these sites to linearize it. A double-stranded oligonucleotide encoding the sequence of the DNA between the restriction sites but containing the desired mutation(s) is synthesized using standard procedures. The two strands are synthesized separately and then hybridized together using standard techniques. This double-stranded oligonucleotide is referred to as the cassette. This cassette is designed to have 3' and 5' ends that are comparable with the ends of the linearized plasmid, such that it can be directly ligated to the plasmid. This plasmid now contains the mutated desired protein subunit DNA sequence.

(D) Combinatorial Mutagenesis

Combinatorial mutagenesis can also be used to generate mutants (Ladner et al., WO 88/06630). In this method, the amino acid sequences for a group of homologs or other related proteins are aligned, preferably to promote the highest homology possible. All of the amino acids which appear at a given position of the aligned sequences can be selected to create a degenerate set of combinatorial sequences. The variegated library of variants is generated by combinatorial mutagenesis at the nucleic acid level, and is encoded by a variegated gene library. For example, a mixture of synthetic oligonucleotides can be enzymatically ligated into gene sequences such that the degenerate set of potential sequences are expressible as individual peptides, or alternatively, as a set of larger fusion proteins containing the set of degenerate sequences.

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Other Modifications of H. pylori Nucleic Acids and Polypeptides

It is possible to modify the structure of an *H. pylori* polypeptide for such purposes as increasing solubility, enhancing stability (e.g., shelf life *ex vivo* and resistance to proteolytic degradation *in vivo*). A modified *H. pylori* protein or peptide can be produced in which the amino acid sequence has been altered, such as by amino acid substitution, deletion, or addition as described herein.

An *H. pylori* peptide can also be modified by substitution of cysteine residues preferably with alanine, serine, threonine, leucine or glutamic acid residues to minimize dimerization via disulfide linkages. In addition, amino acid side chains of fragments of the protein of the invention can be chemically modified. Another modification is cyclization of the peptide.

In order to enhance stability and/or reactivity, an *H. pylori* polypeptide can be modified to incorporate one or more polymorphisms in the amino acid sequence of the protein resulting from any natural allelic variation. Additionally, D-amino acids, nonnatural amino acids, or non-amino acid analogs can be substituted or added to produce a modified protein within the scope of this invention. Furthermore, an *H. pylori* polypeptide can be modified using polyethylene glycol (PEG) according to the method of A. Sehon and co-workers (Wie et al., supra) to produce a protein conjugated with PEG. In addition, PEG can be added during chemical synthesis of the protein. Other modifications of *H. pylori* proteins include reduction/alkylation (Tarr, *Methods of Protein Microcharacterization*, J. E. Silver ed., Humana Press, Clifton NJ 155-194 (1986)); acylation (Tarr, supra); chemical coupling to an appropriate carrier (Mishell and Shiigi, eds, *Selected Methods in Cellular Immunology*, WH Freeman, San Francisco, CA (1980), U.S. Patent 4,939,239; or mild formalin treatment (Marsh, (1971) *Int. Arch. of Allergy and Appl. Immunol.*, 41: 199 - 215).

To facilitate purification and potentially increase solubility of an *H. pylori* protein or peptide, it is possible to add an amino acid fusion moiety to the peptide backbone. For example, hexa-histidine can be added to the protein for purification by immobilized metal ion affinity chromatography (Hochuli, E. et al., (1988) *Bio/Technology*, 6: 1321 - 1325). In addition, to facilitate isolation of peptides free of irrelevant sequences, specific endoprotease cleavage sites can be introduced between the sequences of the fusion moiety and the peptide.

To potentially aid proper antigen processing of epitopes within an *H. pylori* polypeptide, canonical protease sensitive sites can be engineered between regions, each comprising at least one epitope via recombinant or synthetic methods. For example, charged amino acid pairs, such as KK or RR, can be introduced between regions within a protein or fragment during recombinant construction thereof. The resulting peptide

can be rendered sensitive to cleavage by cathepsin and/or other trypsin-like enzymes which would generate portions of the protein containing one or more epitopes. In addition, such charged amino acid residues can result in an increase in the solubility of the peptide.

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Primary Methods for Screening Polypeptides and Analogs

Various techniques are known in the art for screening generated mutant gene products. Techniques for screening large gene libraries often include cloning the gene library into replicable expression vectors, transforming appropriate cells with the resulting library of vectors, and expressing the genes under conditions in which detection of a desired activity, e.g., in this case, binding to *H. pylori* polypeptide or an interacting protein, facilitates relatively easy isolation of the vector encoding the gene whose product was detected. Each of the techniques described below is amenable to high through-put analysis for screening large numbers of sequences created, e.g., by random mutagenesis techniques.

(A) Two Hybrid Systems

Two hybrid assays such as the system described above (as with the other screening methods described herein), can be used to identify polypeptides, e.g., fragments or analogs of a naturally-occurring *H. pylori* polypeptide, e.g., of cellular proteins, or of randomly generated polypeptides which bind to an *H. pylori* protein. (The *H. pylori* domain is used as the bait protein and the library of variants are expressed as fish fusion proteins.) In an analogous fashion, a two hybrid assay (as with the other screening methods described herein), can be used to find polypeptides which bind a *H. pylori* polypeptide.

(B) Display Libraries

In one approach to screening assays, the candidate peptides are displayed on the surface of a cell or viral particle, and the ability of particular cells or viral particles to bind an appropriate receptor protein via the displayed product is detected in a "panning assay". For example, the gene library can be cloned into the gene for a surface membrane protein of a bacterial cell, and the resulting fusion protein detected by panning (Ladner et al., WO 88/06630; Fuchs et al. (1991) *Bio/Technology* 9:1370-1371; and Goward et al. (1992) *TIBS* 18:136-140). In a similar fashion, a detectably labeled ligand can be used to score for potentially functional peptide homologs. Fluorescently labeled ligands, e.g., receptors, can be used to detect homologs which retain ligand-binding activity. The use of fluorescently labeled ligands, allows cells to be visually

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inspected and separated under a fluorescence microscope, or, where the morphology of the cell permits, to be separated by a fluorescence-activated cell sorter.

A gene library can be expressed as a fusion protein on the surface of a viral particle. For instance, in the filamentous phage system, foreign peptide sequences can be expressed on the surface of infectious phage, thereby conferring two significant benefits. First, since these phage can be applied to affinity matrices at concentrations well over 10¹³ phage per milliliter, a large number of phage can be screened at one time. Second, since each infectious phage displays a gene product on its surface, if a particular phage is recovered from an affinity matrix in low yield, the phage can be amplified by another round of infection. The group of almost identical E. coli filamentous phages M13. fd., and f1 are most often used in phage display libraries. Either of the phage gIII or gVIII coat proteins can be used to generate fusion proteins without disrupting the ultimate packaging of the viral particle. Foreign epitopes can be expressed at the NH2terminal end of pIII and phage bearing such epitopes recovered from a large excess of phage lacking this epitope (Ladner et al. PCT publication WO 90/02909; Garrard et al., PCT publication WO 92/09690; Marks et al. (1992) J. Biol. Chem. 267:16007-16010; Griffiths et al. (1993) EMBO J 12:725-734; Clackson et al. (1991) Nature 352:624-628; and Barbas et al. (1992) PNAS 89:4457-4461).

A common approach uses the maltose receptor of E. coli (the outer membrane protein, LamB) as a peptide fusion partner (Charbit et al. (1986) EMBO 5, 3029-3037). Oligonucleotides have been inserted into plasmids encoding the LamB gene to produce peptides fused into one of the extracellular loops of the protein. These peptides are available for binding to ligands, e.g., to antibodies, and can elicit an immune response when the cells are administered to animals. Other cell surface proteins, e.g., OmpA (Schorr et al. (1991) Vaccines 91, pp. 387-392), PhoE (Agterberg, et al. (1990) Gene 88, 37-45), and PAL (Fuchs et al. (1991) Bio/Tech 9, 1369-1372), as well as large bacterial surface structures have served as vehicles for peptide display. Peptides can be fused to pilin, a protein which polymerizes to form the pilus-a conduit for interbacterial exchange of genetic information (Thiry et al. (1989) Appl. Environ. Microbiol. 55, 984-993). Because of its role in interacting with other cells, the pilus provides a useful support for the presentation of peptides to the extracellular environment. Another large surface structure used for peptide display is the bacterial motive organ, the flagellum. Fusion of peptides to the subunit protein flagellin offers a dense array of many peptide copies on the host cells (Kuwajima et al. (1988) Bio/Tech. 6, 1080-1083). Surface proteins of other bacterial species have also served as peptide fusion partners. Examples include the Staphylococcus protein A and the outer membrane IgA protease of Neisseria (Hansson

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et al. (1992) J. Bacteriol. 174, 4239-4245 and Klauser et al. (1990) EMBO J. 9, 1991-1999).

In the filamentous phage systems and the LamB system described above, the physical link between the peptide and its encoding DNA occurs by the containment of the DNA within a particle (cell or phage) that carries the peptide on its surface. Capturing the peptide captures the particle and the DNA within. An alternative scheme uses the DNA-binding protein LacI to form a link between peptide and DNA (Cull et al. (1992) PNAS USA 89:1865-1869). This system uses a plasmid containing the LacI gene with an oligonucleotide cloning site at its 3'-end. Under the controlled induction by arabinose, a LacI-peptide fusion protein is produced. This fusion retains the natural ability of LacI to bind to a short DNA sequence known as LacO operator (LacO). By installing two copies of LacO on the expression plasmid, the LacI-peptide fusion binds tightly to the plasmid that encoded it. Because the plasmids in each cell contain only a single oligonucleotide sequence and each cell expresses only a single peptide sequence, the peptides become specifically and stably associated with the DNA sequence that directed its synthesis. The cells of the library are gently lysed and the peptide-DNA complexes are exposed to a matrix of immobilized receptor to recover the complexes containing active peptides. The associated plasmid DNA is then reintroduced into cells for amplification and DNA sequencing to determine the identity of the peptide ligands. As a demonstration of the practical utility of the method, a large random library of dodecapeptides was made and selected on a monoclonal antibody raised against the opioid peptide dynorphin B. A cohort of peptides was recovered, all related by a consensus sequence corresponding to a six-residue portion of dynorphin B. (Cull et al. (1992) Proc. Natl. Acad. Sci. U.S.A. 89-1869)

This scheme, sometimes referred to as peptides-on-plasmids, differs in two important ways from the phage display methods. First, the peptides are attached to the C-terminus of the fusion protein, resulting in the display of the library members as peptides having free carboxy termini. Both of the filamentous phage coat proteins, pIII and pVIII, are anchored to the phage through their C-termini, and the guest peptides are placed into the outward-extending N-terminal domains. In some designs, the phage-displayed peptides are presented right at the amino terminus of the fusion protein. (Cwirla, et al. (1990) *Proc. Natl. Acad. Sci. U.S.A.* 87, 6378-6382) A second difference is the set of biological biases affecting the population of peptides actually present in the libraries. The LacI fusion molecules are confined to the cytoplasm of the host cells. The phage coat fusions are exposed briefly to the cytoplasm during translation but are rapidly secreted through the inner membrane into the periplasmic compartment, remaining anchored in the membrane by their C-terminal hydrophobic domains, with the

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N-termini, containing the peptides, protruding into the periplasm while awaiting assembly into phage particles. The peptides in the LacI and phage libraries may differ significantly as a result of their exposure to different proteolytic activities. The phage coat proteins require transport across the inner membrane and signal peptidase processing as a prelude to incorporation into phage. Certain peptides exert a deleterious effect on these processes and are underrepresented in the libraries (Gallop et al. (1994) J. Med. Chem. 37(9):1233-1251). These particular biases are not a factor in the LacI display system.

The number of small peptides available in recombinant random libraries is enormous. Libraries of 10⁷-10⁹ independent clones are routinely prepared. Libraries as large as 10¹¹ recombinants have been created, but this size approaches the practical limit for clone libraries. This limitation in library size occurs at the step of transforming the DNA containing randomized segments into the host bacterial cells. To circumvent this limitation, an *in vitro* system based on the display of nascent peptides in polysome complexes has recently been developed. This display library method has the potential of producing libraries 3-6 orders of magnitude larger than the currently available phage/phagemid or plasmid libraries. Furthermore, the construction of the libraries, expression of the peptides, and screening, is done in an entirely cell-free format.

In one application of this method (Gallop et al. (1994) J. Med. Chem. 37(9):1233-1251), a molecular DNA library encoding 1012 decapeptides was constructed and the library expressed in an E. coli S30 in vitro coupled transcription/translation system. Conditions were chosen to stall the ribosomes on the mRNA, causing the accumulation of a substantial proportion of the RNA in polysomes and yielding complexes containing nascent peptides still linked to their encoding RNA. The polysomes are sufficiently robust to be affinity purified on immobilized receptors in much the same way as the more conventional recombinant peptide display libraries are screened. RNA from the bound complexes is recovered, converted to cDNA, and amplified by PCR to produce a template for the next round of synthesis and screening. The polysome display method can be coupled to the phage display system. Following several rounds of screening, cDNA from the enriched pool of polysomes was cloned into a phagemid vector. This vector serves as both a peptide expression vector, displaying peptides fused to the coat proteins, and as a DNA sequencing vector for peptide identification. By expressing the polysome-derived peptides on phage, one can either continue the affinity selection procedure in this format or assay the peptides on individual clones for binding activity in a phage ELISA, or for binding specificity in a completion phage ELISA (Barret, et al. (1992) Anal. Biochem 204,357-364). To

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identify the sequences of the active peptides one sequences the DNA produced by the phagemid host.

Secondary Screening of Polypeptides and Analogs

The high through-put assays described above can be followed by secondary screens in order to identify further biological activities which will, e.g., allow one skilled in the art to differentiate agonists from antagonists. The type of a secondary screen used will depend on the desired activity that needs to be tested. For example, an assay can be developed in which the ability to inhibit an interaction between a protein of interest and its respective ligand can be used to identify antagonists from a group of peptide fragments isolated though one of the primary screens described above.

Therefore, methods for generating fragments and analogs and testing them for activity are known in the art. Once the core sequence of interest is identified, it is routine for one skilled in the art to obtain analogs and fragments.

Peptide Mimetics of H. pylori Polypeptides

The invention also provides for reduction of the protein binding domains of the subject *H. pylori* polypeptides to generate mimetics, e.g. peptide or non-peptide agents. The peptide mimetics are able to disrupt binding of a polypeptide to its counter ligand, e.g., in the case of an *H. pylori* polypeptide binding to a naturally occurring ligand. The critical residues of a subject *H. pylori* polypeptide which are involved in molecular recognition of a polypeptide can be determined and used to generate *H. pylori*-derived peptidomimetics which competitively or noncompetitively inhibit binding of the *H. pylori* polypeptide with an interacting polypeptide (see, for example, European patent applications EP-412,762A and EP-B31,080A).

For example, scanning mutagenesis can be used to map the amino acid residues of a particular *H. pylori* polypeptide involved in binding an interacting polypeptide, peptidomimetic compounds (e.g. diazepine or isoquinoline derivatives) can be generated which mimic those residues in binding to an interacting polypeptide, and which therefore can inhibit binding of an *H. pylori* polypeptide to an interacting polypeptide and thereby interfere with the function of *H. pylori* polypeptide. For instance, non-hydrolyzable peptide analogs of such residues can be generated using benzodiazepine (e.g., see Freidinger et al. in *Peptides: Chemistry and Biology*, G.R. Marshall ed., ESCOM Publisher: Leiden, Netherlands, 1988), azepine (e.g., see Huffman et al. in *Peptides: Chemistry and Biology*, G.R. Marshall ed., ESCOM Publisher: Leiden, Netherlands, 1988), substituted gama lactam rings (Garvey et al. in *Peptides: Chemistry and Biology*, G.R. Marshall ed., ESCOM Publisher: Leiden, Netherlands, 1988), keto-

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methylene pseudopeptides (Ewenson et al. (1986) J Med Chem 29:295; and Ewenson et al. in Peptides: Structure and Function (Proceedings of the 9th American Peptide Symposium) Pierce Chemical Co. Rockland, IL, 1985), β-turn dipeptide cores (Nagai et al. (1985) Tetrahedron Lett 26:647; and Sato et al. (1986) J Chem Soc Perkin Trans 1:1231), and β-aminoalcohols (Gordon et al. (1985) Biochem Biophys Res Commun 126:419; and Dann et al. (1986) Biochem Biophys Res Commun 134:71).

VI. Vaccine Formulations for H. pylori Nucleic Acids and Polypeptides

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This invention also features vaccine compositions or formulations (used interchangeably herein) for protection against infection by H. pylori or for treatment of H. pylori infection. As used herein, the term "treatment of H. pylori infection" refers to therapeutic treatment of an existing or established H. pylori infection. The terms "protection against H. pylori infection" or "prophylactic treatment" refer to the use of H. pylori vaccine formulation for reducing the risk of or preventing an infection in a subject at risk for H. pylori infection. In one embodiment, the vaccine compositions contain one or more immunogenic components, such as a surface protein, from H. pylori, or portion thereof, and a pharmaceutically acceptable carrier. For example, in one embodiment, the vaccine formulations of the invention contain at least one or combination of H. pylori polypeptides or fragments thereof, from same or different H. pylori antigens. Nucleic acids and H. pylori polypeptides for use in the vaccine formulations of the invention include the nucleic acids and polypeptides set forth in the Sequence Listing, preferably those H. pylori nucleic acids that encode surface proteins and surface proteins or fragments thereof. For example, a preferred nucleic acid and H. pylori polypeptide for use in a vaccine composition of the invention is selected from the group of nucleic acids which encode cell envelope proteins and H. pylori cell envelope proteins as set forth in Table 1. However, any nucleic acid encoding an immunogenic H. pylori protein and H. pylori polypetide, or portion thereof, can be used in the present invention. These vaccines have therapeutic and/or prophylactic utilities.

One aspect of the invention provides a vaccine composition for protection against infection by *H. pylori* which contains at least one immunogenic fragment of an *H. pylori* protein and a pharmaceutically acceptable carrier. Preferred fragments include peptides of at least about 10 amino acid residues in length, preferably about 10-20 amino acid residues in length, and more preferably about 12-16 amino acid residues in length.

Immunogenic components of the invention can be obtained, for example, by screening polypeptides recombinantly produced from the corresponding fragment of the nucleic acid encoding the full-length *H. pylori* protein. In addition, fragments can be

chemically synthesized using techniques known in the art such as conventional Merrifield solid phase f-Moc or t-Boc chemistry.

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In one embodiment, immunogenic components are identified by the ability of the peptide to stimulate T cells. Peptides which stimulate T cells, as determined by, for example, T cell proliferation or cytokine secretion are defined herein as comprising at least one T cell epitope. T cell epitopes are believed to be involved in initiation and perpetuation of the immune response to the protein allergen which is responsible for the clinical symptoms of allergy. These T cell epitopes are thought to trigger early events at the level of the T helper cell by binding to an appropriate HLA molecule on the surface of an antigen presenting cell, thereby stimulating the T cell subpopulation with the relevant T cell receptor for the epitope. These events lead to T cell proliferation, lymphokine secretion, local inflammatory reactions, recruitment of additional immune cells to the site of antigen/T cell interaction, and activation of the B cell cascade, leading to the production of antibodies. A T cell epitope is the basic element, or smallest unit of recognition by a T cell receptor, where the epitope comprises amino acids essential to receptor recognition (e.g., approximately 6 or 7 amino acid residues). Amino acid sequences which mimic those of the T cell epitopes are within the scope of this invention.

In another embodiment, immunogenic components of the invention are identified through genomic vaccination. The basic protocol is based on the idea that expression libraries consisting of all or parts of a pathogen genome, e.g., an *H. pylori* genome, can confer protection when used to genetically immunize a host. This expression library immunization (ELI) is analogous to expression cloning and involves reducing a genomic expression library of a pathogen, e.g., *H. pylori*, into plasmids that can act as genetic vaccines. The plasmids can also be designed to encode genetic adjuvants which can dramatically stimulate the humoral response. These genetic adjuvants can be introduced at remote sites and act as well extracelluraly as intracellularly.

This is a new approach to vaccine production that has many of the advantages of live/attenuated pathogens but no risk of infection. An expression library of pathogen DNA is used to immunize a host thereby producing the effects of antigen presentation of a live vaccine without the risk. For example, in the present invention, random fragments from the *H. pylori* genome or from cosmid or plasmid clones, as well as PCR products from genes identified by genomic sequencing, can be used to immunize a host. The feasibility of this approach has been demonstrated with *Mycoplasma pulmonis* (Barry et al., *Nature* 377:632-635, 1995), where even partial expression libraries of *Mycoplasma pulmonis*, a natural pathogen in rodents, provided protection against challenge from the pathogen.

ELI is a technique that allows for production of a non-infectious multipartite vaccine, even when little is known about pathogen's biology, because ELI uses the immune system to screen candidate genes. Once isolated, these genes can be used as genetic vaccines or for development of recombinant protein vaccines. Thus, ELI allows for production of vaccines in a systematic, largely mechanized fashion.

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Screening immunogenic components can be accomplished using one or more of several different assays. For example, *in vitro*, peptide T cell stimulatory activity is assayed by contacting a peptide known or suspected of being immunogenic with an antigen presenting cell which presents appropriate MHC molecules in a T cell culture. Presentation of an immunogenic *H. pylori* peptide in association with appropriate MHC molecules to T cells in conjunction with the necessary costimulation has the effect of transmitting a signal to the T cell that induces the production of increased levels of cytokines, particularly of interleukin-2 and interleukin-4. The culture supernatant can be obtained and assayed for interleukin-2 or other known cytokines. For example, any one of several conventional assays for interleukin-2 can be employed, such as the assay described in *Proc. Natl. Acad. Sci USA*, 86: 1333 (1989) the pertinent portions of which are incorporated herein by reference. A kit for an assay for the production of interferon is also available from Genzyme Corporation (Cambridge, MA).

Alternatively, a common assay for T cell proliferation entails measuring tritiated thymidine incorporation. The proliferation of T cells can be measured *in vitro* by determining the amount of ³H-labeled thymidine incorporated into the replicating DNA of cultured cells. Therefore, the rate of DNA synthesis and, in turn, the rate of cell division can be quantified.

Vaccine compositions or formulations of the invention containing one or more immunogenic components (e.g., H. pylori polypeptide or fragment thereof or nucleic acid encoding an H. pylori polypeptide or fragment thereof) preferably include a pharmaceutically acceptable carrier. The term "pharmaceutically acceptable carrier" is intended to include any and all solvents, dispersion media, coatings, antibacterial and antifungal agents, isotonic and absorption delaying agents, and the like, compatible with pharmaceutical administration. Suitable pharmaceutically acceptable carriers include, for example, one or more of water, saline, phosphate buffered saline, dextrose, glycerol, ethanol and the like, as well as combinations thereof. Pharmaceutically acceptable carriers may further comprise minor amounts of auxiliary substances such as wetting or emulsifying agents, preservatives or buffers, which enhance the shelf life or effectiveness of the H. pylori nucleic acid or polypeptide. For vaccine formulations of the invention containing H. pylori polypeptides, the polypeptide is preferably coadministered with a suitable adjuvant and/or a delivery system described herein.

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It will be apparent to those of skill in the art that the therapeutically effective amount of DNA or protein of this invention will depend, *inter alia*, upon the administration schedule, the unit dose of an *H. pylori* nucleic acid or polypeptide administered, whether the protein or nucleic acid is administered in combination with other therapeutic agents, the immune status and health of the patient, and the therapeutic activity of the particular protein or nucleic acid.

Vaccine formulations are conventionally administered parenterally, e.g., by injection, either subcutaneously or intramuscularly. Methods for intramuscular immunization are described by Wolff et al. (1990) Science 247: 1465-1468 and by Sedegah et al. (1994) Immunology 91: 9866-9870. Other modes of administration include oral and pulmonary formulations, suppositories, and transdermal applications. Oral immunization is preferred over parenteral methods for inducing protection against infection by H. pylori. Czinn et. al. (1993) Vaccine 11: 637-642. Oral formulations include such normally employed excipients as, for example, pharmaceutical grades of mannitol, lactose, starch, magnesium stearate, sodium saccharine, cellulose, magnesium carbonate, and the like.

In one embodiment, the vaccine formulation includes, as a pharmaceutically acceptable carrier, an adjuvant. Examples of the suitable adjuvants for use in the vaccine formulations of the invention include, but are not limited, to aluminum hydroxide; N-acetyl-muramyl-L-threonyl-D-isoglutamine (thr-MDP); N-acetyl-normuramyl-L-alanyl-D-isoglutamine (CGP 11637, referred to as nor-MDP); N-acetylmuramyl-L-alanyl-D-isoglutaminyl-L-alanine-2-(1'-2'-dipalmitoyl-sn-glycero-3-hydroxyphos-phoryloxy)-ethylamine (CGP 19835A, referred to a MTP-PE); RIBI, which contains three components from bacteria; monophosphoryl lipid A; trehalose dimycoloate; cell wall skeleton (MPL + TDM + CWS) in a 2% squalene/Tween 80 emulsion; and cholera toxin. Others which may be used are non-toxic derivatives of cholera toxin, including its B subunit, and/or conjugates or genetically engineered fusions of the *H. pylori* polypeptide with cholera toxin or its B subunit, procholeragenoid, fungal polysaccharides, including schizophyllan, muramyl dipeptide, muramyl dipeptide derivatives, phorbol esters, labile toxin of *E. coli*, non-*H. pylori* bacterial lysates, block polymers or saponins.

In another embodiment, the vaccine formulation includes, as a pharmaceutically acceptable carrier, a delivery system. Suitable delivery systems for use in the vaccine formulations of the invention include biodegradable microcapsules or immunostimulating complexes (ISCOMs), cochleates, or liposomes, genetically engineered attenuated live vectors such as viruses or bacteria, and recombinant (chimeric) virus-like

particles, e.g., bluetongue. In another embodiment of the invention, the vaccine formulation includes both a delivery system and an adjuvant.

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Delivery systems in humans may include enteric release capsules protecting the antigen from the acidic environment of the stomach, and including *H. pylori* polypeptide in an insoluble form as fusion proteins. Suitable carriers for the vaccines of the invention are enteric coated capsules and polylactide-glycolide microspheres. Suitable diluents are 0.2 N NaHCO3 and/or saline.

Vaccines of the invention can be administered as a primary prophylactic agent in adults or in children, as a secondary prevention, after successful eradication of *H. pylori* in an infected host, or as a therapeutic agent in the aim to induce an immune response in a susceptible host to prevent infection by *H. pylori*. The vaccines of the invention are administered in amounts readily determined by persons of ordinary skill in the art. Thus, for adults a suitable dosage will be in the range of 10 µg to 10 g, preferably 10 µg to 100 mg, for example 50 µg to 50 mg. A suitable dosage for adults will also be in the range of 5 µg to 500 mg. Similar dosage ranges will be applicable for children.

The amount of adjuvant employed will depend on the type of adjuvant used. For example, when the mucosal adjuvant is cholera toxin, it is suitably used in an amount of 5 μ g to 50 μ g, for example 10 μ g to 35 μ g. When used in the form of microcapsules, the amount used will depend on the amount employed in the matrix of the microcapsule to achieve the desired dosage. The determination of this amount is within the skill of a person of ordinary skill in the art.

Those skilled in the art will recognize that the optimal dose may be more or less depending upon the patient's body weight, disease, the route of administration, and other factors. Those skilled in the art will also recognize that appropriate dosage levels can be obtained based on results with known oral vaccines such as, for example, a vaccine based on an *E. coli* lysate (6 mg dose daily up to total of 540 mg) and with an enterotoxigenic *E. coli* purified antigen (4 doses of 1 mg) (Schulman et al., *J. Urol.* 150:917-921 (1993)); Boedecker et al., *American Gastroenterological Assoc.* 999:A-222 (1993)). The number of doses will depend upon the disease, the formulation, and efficacy data from clinical trials. Without intending any limitation as to the course of treatment, the treatment can be administered over 3 to 8 doses for a primary immunization schedule over 1 month (Boedeker, *American Gastroenterological Assoc.* 888:A-222 (1993)).

In a preferred embodiment, a vaccine composition of the invention can be based on a killed whole *E. coli* preparation with an immunogenic fragment of an *H. pylori* protein of the invention expressed on its surface or it can be based on an *E. coli* lysate, wherein the killed *E. coli* acts as a carrier or an adjuvant.

It will be apparent to those skilled in the art that some of the vaccine compositions of the invention are useful only for preventing *H. pylori* infection, some are useful only for treating *H. pylori* infection, and some are useful for both preventing and treating *H. pylori* infection. In a preferred embodiment, the vaccine composition of the invention provides protection against *H. pylori* infection by stimulating humoral and/or cell-mediated immunity against *H. pylori*. It should be understood that amelioration of any of the symptoms of *H. pylori* infection is a desirable clinical goal, including a lessening of the dosage of medication used to treat *H. pylori*-caused disease, or an increase in the production of antibodies in the serum or mucous of patients.

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VII. Antibodies Reactive With H. pylori Polypeptides

The invention also includes antibodies specifically reactive with the subject *H. pylori* polypeptide. Anti-protein/anti-peptide antisera or monoclonal antibodies can be made by standard protocols (See, for example, *Antibodies: A Laboratory Manual* ed. by Harlow and Lane (Cold Spring Harbor Press: 1988)). A mammal such as a mouse, a hamster or rabbit can be immunized with an immunogenic form of the peptide. Techniques for conferring immunogenicity on a protein or peptide include conjugation to carriers or other techniques well known in the art. An immunogenic portion of the subject *H. pylori* polypeptide can be administered in the presence of adjuvant. The progress of immunization can be monitored by detection of antibody titers in plasma or serum. Standard ELISA or other immunoassays can be used with the immunogen as antigen to assess the levels of antibodies.

In a preferred embodiment, the subject antibodies are immunospecific for antigenic determinants of the *H. pylori* polypeptides of the invention, e.g. antigenic determinants of a polypeptide of the invention contained in the Sequence Listing, or a closely related human or non-human mammalian homolog (e.g., 90% homologous, more preferably at least 95% homologous). In yet a further preferred embodiment of the invention, the anti-*H. pylori* antibodies do not substantially cross react (i.e., react specifically) with a protein which is for example, less than 80% percent homologous to a sequence of the invention contained in the Sequence Listing. By "not substantially cross react", it is meant that the antibody has a binding affinity for a non-homologous protein which is less than 10 percent, more preferably less than 5 percent, and even more preferably less than 1 percent, of the binding affinity for a protein of the invention contained in the Sequence Listing. In a most preferred embodiment, there is no crossreactivity between bacterial and mammalian antigens.

The term antibody as used herein is intended to include fragments thereof which are also specifically reactive with *H. pylori* polypeptides. Antibodies can be fragmented

using conventional techniques and the fragments screened for utility in the same manner as described above for whole antibodies. For example, F(ab')₂ fragments can be generated by treating antibody with pepsin. The resulting F(ab')₂ fragment can be treated to reduce disulfide bridges to produce Fab' fragments. The antibody of the invention is further intended to include bispecific and chimeric molecules having an anti-H. pylori portion.

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Both monoclonal and polyclonal antibodies (Ab) directed against *H. pylori* polypeptides or *H. pylori* polypeptide variants, and antibody fragments such as Fab' and F(ab')₂, can be used to block the action of *H. pylori* polypeptide and allow the study of the role of a particular *H. pylori* polypeptide of the invention in aberrant or unwanted intracellular signaling, as well as the normal cellular function of the *H. pylori* and by microinjection of anti-*H. pylori* polypeptide antibodies of the present invention.

Antibodies which specifically bind *H. pylori* epitopes can also be used in immunohistochemical staining of tissue samples in order to evaluate the abundance and pattern of expression of *H. pylori* antigens. Anti *H. pylori* polypeptide antibodies can be used diagnostically in immuno-precipitation and immuno-blotting to detect and evaluate *H. pylori* levels in tissue or bodily fluid as part of a clinical testing procedure. Likewise, the ability to monitor *H. pylori* polypeptide levels in an individual can allow determination of the efficacy of a given treatment regimen for an individual afflicted with such a disorder. The level of an *H. pylori* polypeptide can be measured in cells found in bodily fluid, such as in urine samples or can be measured in tissue, such as produced by gastric biopsy. Diagnostic assays using anti-*H. pylori* antibodies can include, for example, immunoassays designed to aid in early diagnosis of *H. pylori* infections. The present invention can also be used as a method of detecting antibodies contained in samples from individuals infected by this bacterium using specific *H. pylori* antigens.

Another application of anti-*H. pylori* polypeptide antibodies of the invention is in the immunological screening of cDNA libraries constructed in expression vectors such as λ gt11, λ gt18-23, λ ZAP, and λ ORF8. Messenger libraries of this type, having coding sequences inserted in the correct reading frame and orientation, can produce fusion proteins. For instance, λ gt11 will produce fusion proteins whose amino termini consist of β -galactosidase amino acid sequences and whose carboxy termini consist of a foreign polypeptide. Antigenic epitopes of a subject *H. pylori* polypeptide can then be detected with antibodies, as, for example, reacting nitrocellulose filters lifted from infected plates with anti-*H. pylori* polypeptide antibodies. Phage, scored by this assay, can then be isolated from the infected plate. Thus, the presence of *H. pylori* gene

homologs can be detected and cloned from other species, and alternate isoforms (including splicing variants) can be detected and cloned.

VIII. Kits Containing Nucleic Acids, Polypeptides or Antibodies of the Invention

The nucleic acid, polypeptides and antibodies of the invention can be combined with other reagents and articles to form kits. Kits for diagnostic purposes typically comprise the nucleic acid, polypeptides or antibodies in vials or other suitable vessels. Kits typically comprise other reagents for performing hybridization reactions, polymerase chain reactions (PCR), or for reconstitution of lyophilized components, such as aqueous media, salts, buffers, and the like. Kits may also comprise reagents for sample processing such as detergents, chaotropic salts and the like. Kits may also comprise immobilization means such as particles, supports, wells, dipsticks and the like. Kits may also comprise labeling means such as dyes, developing reagents, radioisotopes, fluorescent agents, luminescent or chemiluminescent agents, enzymes, intercalating agents and the like. With the nucleic acid and amino acid sequence information provided herein, individuals skilled in art can readily assemble kits to serve their particular purpose. Kits further can include instructions for use.

IX. Drug Screening Assays Using H. pylori Polypeptides

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By making available purified and recombinant *H. pylori* polypeptides, the present invention provides assays which can be used to screen for drugs which are either agonists or antagonists of the normal cellular function, in this case, of the subject *H. pylori* polypeptides, or of their role in intracellular signaling. Such inhibitors or potentiators may be useful as new therapeutic agents to combat *H. pylori* infections in humans. A variety of assay formats will suffice and, in light of the present inventions, will be comprehended by the skilled artisan.

In many drug screening programs which test libraries of compounds and natural extracts, high throughput assays are desirable in order to maximize the number of compounds surveyed in a given period of time. Assays which are performed in cell-free systems, such as may be derived with purified or semi-purified proteins, are often preferred as "primary" screens in that they can be generated to permit rapid development and relatively easy detection of an alteration in a molecular target which is mediated by a test compound. Moreover, the effects of cellular toxicity and/or bioavailability of the test compound can be generally ignored in the *in vitro* system, the assay instead being focused primarily on the effect of the drug on the molecular target as may be manifest in an alteration of binding affinity with other proteins or change in enzymatic properties of the molecular target. Accordingly, in an exemplary screening assay of the present

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invention, the compound of interest is contacted with an isolated and purified *H. pylori* polypeptide.

Screening assays can be constructed *in vitro* with a purified *H. pylori* polypeptide or fragment thereof, such as an *H. pylori* polypeptide having enzymatic activity, such that the activity of the polypeptide produces a detectable reaction product. The efficacy of the compound can be assessed by generating dose response curves from data obtained using various concentrations of the test compound. Moreover, a control assay can also be performed to provide a baseline for comparison. Suitable products include those with distinctive absorption, fluorescence, or chemi-luminescence properties, for example, because detection may be easily automated. A variety of synthetic or naturally occurring compounds can be tested in the assay to identify those which inhibit or potentiate the activity of the *H. pylori* polypeptide. Some of these active compounds may directly, or with chemical alterations to promote membrane permeability or solubility, also inhibit or potentiate the same activity (e.g., enzymatic activity) in whole, live *H. pylori* cells.

This invention is further illustrated by the following examples which should not be construed as limiting. The contents of all references and published patent applications cited throughout this application are hereby incorporated by reference.

EXEMPLIFICATION

I. Cloning and Sequencing of H. pylori DNA

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H. pylori chromosomal DNA was isolated according to a basic DNA protocol outlined in Schleif R.F. and Wensink P.C., Practical Methods in Molecular Biology, p.98, Springer-Verlag, NY., 1981, with minor modifications. Briefly, cells were pelleted, resuspended in TE (10 mM Tris, 1 mM EDTA, pH 7.6) and GES lysis buffer (5.1 M guanidium thiocyanate, 0.1 M EDTA, pH 8.0, 0.5% N-laurylsarcosine) was added. Suspension was chilled and ammonium acetate (NH₄Ac) was added to final concentration of 2.0 M. DNA was extracted, first with chloroform, then with phenol-chloroform, and reextracted with chloroform. DNA was precipitated with isopropanol, washed twice with 70% EtOH, dried and resuspended in TE.

Following isolation whole genomic *H. pylori* DNA was nebulized (Bodenteich et al., *Automated DNA Sequencing and Analysis* (J.C. Venter, ed.), Academic Press, 1994) to a median size of 2000 bp. After nebulization, the DNA was concentrated and separated on a standard 1% agarose gel. Several fractions, corresponding to

approximate sizes 900-1300 bp, 1300-1700 bp, 1700-2200 bp, 2200-2700 bp, were excised from the gel and purified by the GeneClean procedure (Bio101, Inc.).

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The purified DNA fragments were then blunt-ended using T4 DNA polymerase. The healed DNA was then ligated to unique BstXI-linker adapters in 100-1000 fold molar excess. These linkers are complimentary to the BstXI-cut pMPX vectors, while the overhang is not self-complimentary. Therefore, the linkers will not concatemerize nor will the cut-vector religate itself easily. The linker-adopted inserts were separated from the unincorporated linkers on a 1% agarose gel and purified using GeneClean. The linker-adopted inserts were then ligated to each of the 20 pMPX vectors to construct a series of "shotgun" subclone libraries. The vectors contain an out-of-frame lacZ gene at the cloning site which becomes in-frame in the event that an adapter-dimer is cloned, allowing these to be avoided by their blue-color.

All subsequent steps were based on the multiplex DNA sequencing protocols outlined in Church G.M. and Kieffer-Higgins S., *Science* 240:185-188, 1988. Only major modifications to the protocols are highlighted. Briefly, each of the 20 vectors was then transformed into DH5α competent cells (Gibco/BRL, DH5α transformation protocol). The libraries were assessed by plating onto antibiotic plates containing ampicillin, methicillin and IPTG/Xgal. The plates were incubated overnight at 37°C. Successful transformants were then used for plating of clones and pooling into the multiplex pools. The clones were picked and pooled into 40 ml growth medium cultures. The cultures were grown overnight at 37°C. DNA was purified using the Qiagen Midi-prep kits and Tip-100 columns (Qiagen, Inc.). In this manner, 100 μg of DNA was obtained per pool. Fifteen 96-well plates of DNA were generated to obtain a 5-10 fold sequence redundancy assuming 250-300 base average read-lengths.

These purified DNA samples were then sequenced using the multiplex DNA sequencing based on chemical degradation methods (Church G.M. and Kieffer-Higgins S., Science 240:185-188, 1988) or by Sequithrem (Epicenter Technologies) dideoxy sequencing protocols. The sequencing reactions were electrophoresed and transferred onto nylon membranes by direct transfer electrophoresis from 40 cm gels (Richterich P. and Church G.M., Methods in Enzymology 218:187-222, 1993) or by electroblotting (Church, supra). 24 samples were run per gel. 45 successful membranes were produced by chemical sequencing and 8 were produced by dideoxy sequencing. The DNA was covalently bound to the membranes by exposure to ultraviolet light, and hybridized with labeled oligonucleotides complimentary to tag sequences on the vectors (Church, supra). The membranes were washed to rinse off non-specifically bound probe, and exposed to X-ray film to visualize individual sequence ladders. After autoradiography, the hybridized probe was removed by incubation at 65° C, and the hybridization cycle

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repeated with another tag sequence until the membrane had been probed 38 times for chemical sequencing membranes and 10 times for the dideoxy sequencing membranes. Thus, each gel produced a large number of films, each containing new sequencing information. Whenever a new blot was processed, it was initially probed for an internal standard sequence added to each of the pools.

Digital images of the films were generated using a laser-scanning densitometer (Molecular Dynamics, Sunnyvale, CA). The digitized images were processed on computer workstations (VaxStation 4000's) using the program REPLICA™ (Church et al., Automated DNA Sequencing and Analysis (J.C. Venter, ed.), Academic Press, 1994). Image processing included lane straightening, contrast adjustment to smooth out intensity differences, and resolution enhancement by iterative gaussian deconvolution. The sequences were then automatically picked in REPLICA™ and displayed for interactive proofreading before being stored in a project database. The proofreading was accomplished by a quick visual scan of the film image followed by mouse clicks on the bands of the displayed image to modify the base calls. Many of the sequence errors could be detected and corrected because multiple sequence reads covering the same portion of the genomic DNA provide adequate sequence redundancy for editing. Each sequence automatically received an identification number (corresponding to microtiter plate, probe information, and lane set number). This number serves as a permanent identifier of the sequence so it is always possible to identify the original of any particular sequence without recourse to a specialized database.

Routine assembly of *H. pylori* sequences was done using the program FALCON (Church, Church et al., *Automated DNA Sequenicng and Analysis* (J.C. Venter, ed.), Academic Press, 1994). This program has proven to be fast and reliable for most sequences. The assembled contigs were displayed using a modified version of GelAssemble, developed by the Genetics Computer Group (GCG) (Devereux et al., *Nucleic Acid Res.* 12:387-95, 1984) that interacts with REPLICATM. This provided for an integrated editor that allows multiple sequence gel images to be instantaneously called up from the REPLICATM database and displayed to allow rapid scanning of contigs and proofreading of gel traces where discrepancies occurred between different sequence reads in the assembly.

II. Identification, cloning and expression of recombinant H. pylori DNA sequences

To facilitate the cloning, expression and purification of membrane and secreted proteins from *H. pylori* a powerful gene expression system, the pET System (Novagen), for cloning and expression of recombinant proteins in *E. coli*, was selected. Also, a DNA sequence encoding a peptide tag, the His-Tag, was fused to the 3' end of DNA

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sequences of interest in order to facilitate purification of the recombinant protein products. The 3' end was selected for fusion in order to avoid alteration of any 5' terminal signal sequence. The exception to the above was ppiB, a gene cloned for use as a control in the expression studies. In this study, the sequence for *H. pylori* ppiB contains a DNA sequence encoding a His-Tag fused to the 5' end of the full length gene, because the protein product of this gene does not contain a signal sequence and is expressed as a cytosolic protein.

PCR Amplification and cloning of DNA sequences containing ORF's for membrane and secreted proteins from the J99 Strain of Helicobacter pylori.

Sequences chosen (from the list of the DNA sequences of the invention) for cloning from the J99 strain of H. pylori were prepared for amplification cloning by polymerase chain reaction (PCR). Synthetic oligonucleotide primers (Table 3) specific for the 5' and 3' ends of open reading frames (ORFs) were designed and purchased (GibcoBRL Life Technologies, Gaithersburg, MD, USA). All forward primers (specific for the 5' end of the sequence) were designed to include an Ncol cloning site at the extreme 5' terminus, except for HpSeq. 4821082 where Ndel was used. These primers were designed to permit initiation of protein translation at a methionine residue followed by a valine residue and the coding sequence for the remainder of the native H. pylori DNA sequence. An exception is H. pylori sequence 4821082 where the initiator methionine is immediately followed by the remainder of the native H. pylori DNA sequence. All reverse primers (specific for the 3' end of any H. pylori ORF) included a EcoRI site at the extreme 5' terminus to permit cloning of each H. pylori sequence into the reading frame of the pET-28b. The pET-28b vector provides sequence encoding an additional 20 carboxy-terminal amino acids (only 19 amino acids in HpSeq. 26380318 and HpSeq.14640637) including six histidine residues (at the extreme C-terminus), which comprise the His-Tag. An exception to the above, as noted earlier, is the vector construction for the ppiB gene. A synthetic oligonucleotide primer specific for the 5' end of ppiB gene encoded a BamHI site at its extreme 5' terminus and the primer for the 3' end of the ppiB gene encoded a XhoI site at its extreme 5' terminus.

<u>TABLE 3</u>
<u>Oligonucleotide primers used for PCR amplification of *H. pylori* DNA sequences</u>

Outer membrane Proteins	Forward primer 5' to 3'	Reverse Primer 5' to 3'
Protein 16225006	5'-TATACCATGGTGGG CGCTAA-3' (SEQ ID NO:147)	5'- ATGAATTCGAGTAAG GATTTTTG-3' (SEQ ID NO:148)
Protein 26054702	5'- TTAACCATGGTGAAA AGCGATA-3' (SEQ ID NO:149)	5'- TAGAATTCGCATAAC GATCAATC-3' (SEQ ID NO:150)
Protein 7116626	5'- ATATCCATGGTGAGT TTGATGA-3' (SEQ ID NO:151)	5'- ATGAATTCAATTTT TATTTTGCCA-3' (SEQ ID NO:152)
Protein 29479681	5'- AATTCCATGGTGGGG GCTATG-3' (SEQ ID NO:153)	5'- ATGAATTCTCGATAG CCAAAATC-3' (SEQ ID NO:154)
Protein 14640637	5'- AATTCCATGGTGCAT AACTTCCATT-3' (SEQ ID NO:155)	5'- AAGAATTCTCTAGCA TCCAAATGGA-3' (SEQ ID NO:156)
Periplasmic/ Secreted Proteins		
Protein 30100332	5'-ATTTCCATGGTCATG TCTCATATT-3' (SEQ ID NO:157)	5'- ATGAATTCCATCTTT TATTCCAC-3' (SEQ ID NO:158)
Protein 4721061	5'-AACCATGGTGATTT TAAGCATTGAAAG-3' (SEQ ID NO:159)	5'- AAGAATTCCACTCA AAATTTTTTAACAG-3' (SEQ ID NO:160)
Other Surface Proteins		
Protein 4821082	5'-GATCATCCATATGTT ATCTTCTAAT-3' (SEQ ID NO:161)	5'- TGAATTCAACCATTT TAACCCTG-3' (SEQ ID NO:162)

Protein 978477	5'-TATACCATGGTGAA ATTTTTTCTTTTA-3' (SEQ ID NO:163)	5'- AGAATTCAATTGCG TCTTGTAAAAG-3' (SEQ ID NO:164)
Inner Membrane		
Protein		
		SLATICA ATTOCCA CTT
Protein 26380318	5'-TATACCATGGTGAT	5'-ATGAATTCCCACTT
	GGACAAACTC-3' (SEQ	GGGGCGATA-3' (SEQ ID NO:166)
	ID NO:165)	ID NO.100)
Cytoplasmic Protein		
	5'-TTATGGATCCAAAC	5'-TATCTCGAGTTATA
ppi	CAATTAAAACT-3' (SEQ	GAGAAGGGC-3' (SEQ
	ID NO:167)	ID NO:168)
	110 110.107)	

Genomic DNA prepared from the J99 strain of *H. pylori* (ATCC #55679; deposited by Genome Therapeutics Corporation, 100 Beaver Street, Waltham, MA 02154) was used as the source of template DNA for PCR amplification reactions (Current Protocols in Molecular Biology, John Wiley and Sons, Inc., F. Ausubel et al., eds., 1994). To amplify a DNA sequence containing an *H. pylori* ORF, genomic DNA (50 nanograms) was introduced into a reaction vial containing 2 mM MgCl₂, 1 micromolar synthetic oligonucleotide primers (forward and reverse primers) complementary to and flanking a defined *H. pylori* ORF, 0.2 mM of each deoxynucleotide triphosphate; dATP, dGTP, dCTP, dTTP and 2.5 units of heat stable DNA polymerase (Amplitaq, Roche Molecular Systems. Inc., Branchburg, NJ, USA) in a final volume of 100 microliters. The following thermal cycling conditions were used to obtain amplified DNA products for each ORF using a Perkin Elmer Cetus/ GeneAmp PCR System 9600 thermal cycler:

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Protein 26054702, Protein 7116626, Protein 29479681, Protein 30100332, and Protein 4821082;

Denaturation at 94°C for 2 min,

2 cycles at 94°C for 15 sec, 30°C for 15 sec and 72°C for 1.5 min

23 cycles at 94°C for 15 sec, 55°C for 15 sec and 72°C for 1.5 min

Reactions were concluded at 72°C for 6 minutes.

Protein 16225006;

Denaturation at 94°C for 2 min.

25 cycles at 95°C for 15 sec, 55°C for 15 sec and 72°C for 1.5 min Reaction was concluded at 72°C for 6 minutes.

Protein 4721061;

Denaturation at 94°C for 2 min, 2 cycles at 94°C for 15 sec, 36°C for 15 sec and 72°C for 1.5 min 23 cycles at 94°C for 15 sec, 60°C for 15 sec and 72°C for 1.5 min Reactions were concluded at 72°C for 6 minutes.

Protein 26380318;

Denaturation at 94°C for 2 min, 2 cycles at 94°C for 15 sec, 38°C for 15 sec and 72°C for 1.5 min 23 cycles at 94°C for 15 sec, 62°C for 15 sec and 72°C for 1.5 min Reactions were concluded at 72°C for 6 minutes.

Protein 14640637;

Denaturation at 94°C for 2 min, 2 cycles at 94°C for 15 sec, 33°C for 15 sec and 72°C for 1.5 min 30 cycles at 94°C for 15 sec, 55°C for 15 sec and 72°C for 1.5 min Reactions were concluded at 72°C for 6 minutes.

Conditions for amplification of H. pylori ppiB;

Denaturation at 94°C for 2 min.

2 cycles at 94°C for 15 sec, 32°C for 15 sec and 72°C for 1.5 min

25 cycles at 94°C for 15 sec, 56°C for 15 sec and 72°C for 1.5 min

Reactions were concluded at 72°C for 6 minutes

Upon completion of thermal cycling reactions, each sample of amplified DNA was washed and purified using the Qiaquick Spin PCR purification kit (Qiagen, Gaithersburg, MD, USA). All amplified DNA samples were subjected to digestion with the restriction endonucleases, NcoI and EcoRI (New England BioLabs, Beverly, MA, USA), or in the case of HpSeq. 4821082 (SEQ ID NO: 1309), with NdeI and EcoRI (Current Protocols in Molecular Biology, John Wiley and Sons, Inc., F. Ausubel et al., eds., 1994). DNA samples were then subjected to electrophoresis on 1.0 % NuSeive (FMC BioProducts, Rockland, ME USA) agarose gels. DNA was visualized by exposure to ethidium bromide and long wave uv irradiation. DNA contained in slices

isolated from the agarose gel was purified using the Bio 101 GeneClean Kit protocol (Bio 101 Vista, CA, USA).

Cloning of H. pylori DNA sequences into the pET-28b prokaryotic expression vector.

The pET-28b vector was prepared for cloning by digestion with Ncol and EcoRI, or in the case of *H. pylori* protein 4821082 with Ndel and EcoRI (Current Protocols in Molecular Biology, John Wiley and Sons, Inc., F. Ausubel et al., eds., 1994). In the case of cloning ppiB, the pET-28a vector, which encodes a His-Tag that can be fused to the 5' end of an inserted gene, was used and the cloning site prepared for cloning with the ppiB gene by digestion with BamHI and XhoI restriction endonucleases.

Following digestion, DNA inserts were cloned (Current Protocols in Molecular Biology, John Wiley and Sons, Inc., F. Ausubel et al., eds., 1994) into the previously digested pET-28b expression vector, except for the amplified insert for ppiB, which was cloned into the pET-28a expression vector. Products of the ligation reaction were then used to transform the BL21 strain of *E. coli* (Current Protocols in Molecular Biology, John Wiley and Sons, Inc., F. Ausubel et al., eds., 1994) as described below.

Transformation of competent bacteria with recombinant plasmids

Competent bacteria, *E coli* strain BL21 or *E. coli* strain BL21(DE3), were transformed with recombinant pET expression plasmids carrying the cloned *H. pylori* sequences according to standard methods (Current Protocols in Molecular, John Wiley and Sons, Inc., F. Ausubel et al., eds., 1994). Briefly, 1 microliter of ligation reaction was mixed with 50 microliters of electrocompetent cells and subjected to a high voltage pulse, after which, samples were incubated in 0.45 milliliters SOC medium (0.5% yeast extract, 2.0 % tryptone, 10 mM NaCl, 2.5 mM KCl, 10 mM MgCl2, 10 mM MgSO4 and 20, mM glucose) at 37°C with shaking for 1 hour. Samples were then spread on LB agar plates containing 25 microgram/ml kanamycin sulfate for growth overnight. Transformed colonies of BL21 were then picked and analyzed to evaluate cloned inserts as described below.

Identification of recombinant pET expression plasmids carrying H. pylori sequences

Individual BL21 clones transformed with recombinant pET-28b-H.pylori ORFs were analyzed by PCR amplification of the cloned inserts using the same forward and reverse primers, specific for each *H. pylori* sequence, that were used in the original PCR amplification cloning reactions. Successful amplification verified the integration of the *H. pylori* sequences in the expression vector (Current Protocols in Molecular Biology, John Wiley and Sons, Inc., F. Ausubel et al., eds., 1994).

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Isolation and Preparation of plasmid DNA from BL21 transformants

Individual clones of recombinant pET-28b vectors carrying properly cloned *H. pylori* ORFs were picked and incubated in 5 mls of LB broth plus 25 microgram/ml kanamycin sulfate overnight. The following day plasmid DNA was isolated and purified using the Qiagen plasmid purification protocol (Qiagen Inc., Chatsworth, CA, USA).

Expression of recombinant H. pylori sequences in E. coli

The pET vector can be propagated in any *E. coli* K-12 strain e.g. HMS174, HB101, JM109, DH5, etc. for the purpose of cloning or plasmid preparation. Hosts for expression include *E. coli* strains containing a chromosomal copy of the gene for T7 RNA polymerase. These hosts are lysogens of bacteriophage DE3, a lambda derivative that carries the lacI gene, the lacUV5 promoter and the gene for T7 RNA polymerase. T7 RNA polymerase is induced by addition of isopropyl-B-D-thiogalactoside (IPTG), and the T7 RNA polymerase transcribes any target plasmid, such as pET-28b, carrying a T7 promoter and a gene of interest. Strains used include: BL21(DE3) (Studier, F.W., Rosenberg, A.H., Dunn, J.J., and Dubendorff, J.W. (1990) Meth. Enzymol. 185, 60-89).

To express recombinant *H. pylori* sequences, 50 nanograms of plasmid DNA isolated as described above was used to transform competent BL21(DE3) bacteria as described above (provided by Novagen as part of the pET expression system kit). The lacZ gene (beta-galactosidase) was expressed in the pET-System as described for the *H. pylori* recombinant constructions. Transformed cells were cultured in SOC medium for 1 hour, and the culture was then plated on LB plates containing 25 micrograms/ml kanamycin sulfate. The following day, bacterial colonies were pooled and grown in LB medium containing kanamycin sulfate (25 micrograms/ml) to an optical density at 600 nM of 0.5 to 1.0 O.D. units, at which point, 1 millimolar IPTG was added to the culture for 3 hours to induce gene expression of the *H. pylori* recombinant DNA constructions.

After induction of gene expression with IPTG, bacteria were pelleted by centrifugation in a Sorvall RC-3B centrifuge at 3500 x g for 15 minutes at 4°C. Pellets were resuspended in 50 milliliters of cold 10 mM Tris-HCl, pH 8.0, 0.1 M NaCl and 0.1 mM EDTA (STE buffer). Cells were then centrifuged at 2000 x g for 20 min at 4°C. Wet pellets were weighed and frozen at -80°C until ready for protein purification.

III. Purification of recombinant proteins from E. coli Analytical Methods

The concentrations of purified protein preparations were quantified spectrophotometrically using absorbance coefficients calculated from amino acid

content (Perkins, S.J. 1986 Eur. J. Biochem. 157, 169-180). Protein concentrations were also measured by the method of Bradford, M.M. (1976) Anal. Biochem. 72, 248-254, and Lowry, O.H., Rosebrough, N., Farr, A.L. & Randall, R.J. (1951) J. Biol. Chem. 193, pages 265-275, using bovine serum albumin as a standard.

SDS-polyacrylamide gels (12% or 4.0 to 25 % acrylamide gradient gels) were purchased from BioRad (Hercules, CA, USA), and stained with Coomassie blue. Molecular weight markers included rabbit skeletal muscle myosin (200 kDa), *E. coli* (galactosidase (116 kDa), rabbit muscle phosphorylase B (97.4 kDa), bovine serum albumin (66.2 kDa), ovalbumin (45 kDa), bovine carbonic anhydrase (31 kDa), soybean trypsin inhibitor (21.5 kDa), egg white lysozyme (14.4 kDa) and bovine aprotinin (6.5 kDa).

1. Purification of soluble proteins

All steps were carried out at 4°C. Frozen cells were thawed, resuspended in 5 volumes of lysis buffer (20 mM Tris, pH 7.9, 0.5 M NaCl, 5 mM imidazole with 10% glycerol, 0.1 % 2-mercaptoethanol, 200 µg/ ml lysozyme, 1 mM phenylmethylsulfonyl fluoride (PMSF), and 10 ug/ml each of leupeptin, aprotinin, pepstatin, L-1-chloro-3-[4-tosylamido]-7-amino-2-heptanone (TLCK), L-1-chloro-3-[4-tosylamido]-4-phenyl-2-butanone (TPCK), and soybean trypsin inhibitor, and ruptured by several passages through a small volume microfluidizer (Model M-110S, Microfluidics International Corporation, Newton, MA). The resultant homogenate was made 0.1 % Brij 35, and centrifuged at 100,000 x g for 1 hour to yield a clear supernatant (crude extract).

Following filtration through a 0.8 µm Supor filter (Gelman Sciences, FRG) the crude extract was loaded directly onto a Ni²⁺⁻ nitrilotriacetate-agarose (NTA) with a 5 milliliter bed volume (Hochuli, E., Dbeli, H., and Schacheer, A. (1987) J. Chromatography 411, 177-184) pre-equilibrated in lysis buffer containing 10 % glycerol, 0.1 % Brij 35 and 1 mM PMSF. The column was washed with 250 ml (50 bed volumes) of lysis buffer containing 10 % glycerol, 0.1 % Brij 35, and was eluted with sequential steps of lysis buffer containing 10 % glycerol, 0.05 % Brij 35, 1 mM PMSF, and 20, 100, 200, and 500 mM imidazole in succession. Fractions were monitored by absorbance at OD₂₈₀ nm, and peak fractions were analyzed by SDS-PAGE. Fractions containing the recombinant protein eluted at 100 mM imidazole.

Recombinant protein 14640637 and proteins, beta-galactosidase (lacZ) and peptidyl-prolyl cis-trans isomerase (ppiB)

Fractions containing the recombinant proteins from the Ni²⁺-NTA-agarose columns were pooled and then concentrated to approximately 5 ml by centrifugal

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filtration (Centriprep-10, Amicon, MA), and loaded directly onto a 180-ml column (1.6 X 91 cm) of Sephacryl S-100 HR gel filtration medium equilibrated in Buffer A (10 mM Hepes, pH 7.5, 150 mM NaCl, 0.1 mM EGTA) and run in Buffer A at 18 ml/h. Fractions containing the recombinant protein were identified by absorbance at 280 nm and analyzed by SDS-PAGE. Fractions were pooled and concentrated by centrifugal filtration.

Recombinant protein 7116626

Fractions containing the recombinant protein from the Ni²⁺-NTA-agarose column were pooled and dialyzed overnight against 1 liter of dialysis buffer (10 mM MOPS, pH 6.5, 50 mM NaCl, 0.1 mM EGTA, 0.02% Brij 35 and 1 mM PMSF). In the morning, a fine white precipitate was removed by centrifugation and the resulting supernatant was loaded onto an 8 ml (8 x 75 mm) MonoS high performance liquid chromatography column (Pharmacia Biotechnology, Inc., Piscataway, NJ, USA) equilibrated in buffer B (10 mM MOPS, pH 6.5, 0.1 mM EGTA) containing 50 mM NaCl. The column was washed with 10 bed volumes of buffer B containing 50 mM NaCl, and developed with a 50-ml linear gradient of increasing NaCl (50 to 500 mM). Recombinant protein 7116626 eluted as a sharp peak at 300 mM NaCl.

2. Purification of insoluble proteins from inclusion bodies

The following steps were carried out at 4°C. Cell pellets were resuspended in lysis buffer with 10% glycerol 200 µg/ ml lysozyme, 5 mM EDTA, 1mM PMSF and 0.1% -mercaptoethanol. After passage through the cell disrupter, the resulting homogenate was made 0.2% deoxycholate, stirred 10 minutes, then centrifuged at 20,000 x g, for 30 min. The pellets were washed with lysis buffer containing 10% glycerol, 10 mM EDTA, 1% Triton X-100, 1 mM PMSF and 0.1% -mercaptoethanol, followed by several washes with lysis buffer containing 1 M urea, 1 mM PMSF and 0.1% 2-mercaptoethanol. The resulting white pellet was composed primarily of inclusion bodies, free of unbroken cells and membranous materials..

Recombinant proteins 26054702, 16225006, 30100332, 4721061

The following steps were carried out at room temperature. Purified inclusion bodies were dissolved in 20 ml 8.0 M urea in lysis buffer with 1 mM PMSF and 0.1 % 2-mercaptoethanol, and incubated at room temperature for 1 hour. Materials that did not dissolve were removed by centrifugation. The clear supernatant was filtered, then loaded onto a Ni²⁺-NTA agarose column pre-equilibrated in 8.0 M urea in Lysis Buffer. The column was washed with 250 ml (50 bed volumes) of lysis buffer

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containing 8 M urea, 1.0 mM PMSF and 0.1 % 2-mercaptoethanol, and developed with sequential steps of lysis buffer containing 8M urea, 1 mM PMSF, 0.1 % 2-mercaptoethanol and 20, 100, 200, and 500 mM imidazole in succession. Fractions were monitored by absorbance at OD_{280} nm, and peak fractions were analyzed by SDS-PAGE. Fractions containing the recombinant protein eluted at 100 mM imidazole.

Recombinant proteins 29479681, 26380318

The pellet containing the inclusion bodies was solubilized in buffer B containing 8 M urea, 1 mM PMSF and 0.1 % 2-mercaptoethanol, and incubated for 1 hour at room temperature. Insoluble materials were removed by centrifugation at 20,000 x g for 30 min, and the cleared supernatant was loaded onto a 15 ml (1.6 x 7.5 cm) SP-Sepharose column pre-equilibrated in buffer B, 6 M urea, 1 mM PMSF, 0.1 % 2-mercaptoethanol. After washing the column with 10 bed volumes, the column was developed with a linear gradient from 0 to 500 mM NaCl.

Dialysis and concentration of protein samples

Urea was removed slowly from the protein samples by dialysis against Trisbuffered saline (TBS; 10 mM Tris pH 8.0, 150 mM NaCl) containing 0.5 % deoxycholate (DOC) with sequential reduction in urea concentration as follows; 6M, 4M, 3M, 2M, 1M, 0.5 M and finally TBS without any urea. Each dialysis step was conducted for a minimum of 4 hours at room temperature.

After dialysis, samples were concentrated by pressure filtration using Amicon stirred-cells. Protein concentrations were measured using the methods of Perkins (1986 Eur. J. Biochem. 157, 169-180), Bradford ((1976) Anal. Biochem. 72, 248-254) and Lowry ((1951) J. Biol. Chem. 193, pages 265-275).

The recombinant proteins purified by the methods described above are summarized in Table 4 below.

TABLE 4

J99	Homolog	Gene	Bacterial cell	Method of	Relative	Final	Composit
Sequence Identifier	identified by Blast	symbol of	fraction used to	purification		concentration of purified	
identifier	by Blast	Homolog	, ,		PAGE gel	protein	
			proteins				

Outer Membrane Proteins

16005006	D28626	VEAC	Inclusion bodies	Uia Tag	18 kDa	5 mg/ml	B
16225006	P28635	YEAC	inclusion bodies	HIS- I ag	10 KD4	J mg/m	
26054702	P15929	flgH	Inclusion bodies	His-Tag	37 kDa	1.18 mg/ml	В

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							as dry pellet
7116626	P26093	e(P4)	Soluble fraction	His-Tag	29 kDa	0.8 mg/ml	A
		 		ļ	ļ	1.85 mg/ml	С
29479681	P13036	fecA	Inclusions bodies	SP- Sepharose	23 kDa	2.36 mg/ml	В
						0.5 mg ml	В
							as dry pellet
14640637	P16665	TPF1	Soluble fraction	His-Tag	17 kDa	2.4 mg/ml	A
					ion S100 HR		
Periplasmic/	Secreted Pro	tein	1			1 1	
3010032	P23847	dppA	Inclusion bodies	His-Tag	11 kDa	2.88 mg/mi	В
4721061	P36175	GCP	Inclusion bodies	His-Tag	38 kDa	2.8 mg/mi	В
041 - 5 - 6	- D4-in-	<u> </u>	<u> </u>		L	<u> </u>	
Other Surfac	e Proteins	T	T		<u> </u>	1	
4821082	P08089	M protein	Inclusion bodies	His-Tag	20 kDa	1.16 mg/ml	В
978477	L28919	FBP54	Inclusion bodies	SP-	44 kDa	2.56 mg/ml	В
770477	LZG	1 1 1 3 4	inclusion bodies	Sepharose	44 804	2.30	
						0.3 mg/ml	В
Inner Memb	rane Proteins	s T				· · · · · · · · · · · · · · · · · · ·	
26380318	P15933	ЯiG	Inclusion bodies	SP- Sepharose	11 kDa	22 mg/ml	В
Control Prote	eins with His	-Tag					
	200500		0.111.6				
	P00722	lacZ	Soluble fraction	His-Tag	116 kDa on S200 HR	10 mg/mi	A
-				501 Hillall	OII OZOU FIR	 	
 		рріВ	Soluble fraction	His-Tag	21 kDa	4.4 mg/ml	A
	- 1				on S100 HR		
Buffer composition							
s: A=10 mM He	pes pH 7.5. 1	50 mM Na	Cl, 0.1 mM EGTA			 	
B= 10 mM Tr						1	
			aCl, 0.1 EGTA				

IV. Analysis of H. pylori proteins as Vaccine candidates

To investigate the immunomodulatory effect of *H. pylori* proteins, a mouse/*H. pylori* model was used. This model mimics the human *H. pylori* infection in many respects. The focus is on the effect of oral immunization in *H. pylori* infected animals in order to test the concept of therapeutic oral immunotherapy.

Animals

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Female SPF BALB/c mice were purchased from Bomholt Breeding center (Denmark). They were kept in ordinary makrolon cages with free supply of water and food. The animals were 4-6 weeks old at arrival.

Infection

After a minimum of one week of acclimatization, the animals were infected with a type 2 strain (VacA negative) of *H. pylori* (strain 244, originally isolated from an ulcer patient). In our hands, this strain has earlier proven to be a good colonizer of the mouse stomach. The bacteria were grown overnight in Brucella broth supplemented with 10 % fetal calf serum, at 37°C in a microaerophilic atmosphere (10% CO₂, 5%O₂). The animals were given an oral dose of omeprazole (400 µmol/kg) and 3-5 h after this an oral inoculation of *H. pylori* in broth (approximately 10⁸ cfu/animal). Positive take of the infection was checked in some animals 2-3 weeks after the inoculation.

Antigens

Recombinant *H. pylori* antigens were chosen based on their association with externally exposed *H. pylori* cell membrane. These antigens were selected from the following groups: (1.) Outer Membrane Proteins; (2.) Periplastic/Secreted proteins; (3.) Outer Surface proteins; and (4.) Inner Membrane proteins. All recombinant proteins were constructed with a hexa-HIS tag for purification reasons and the non-*Helicobacter pylori* control protein (b-galactosidase from *E. coli*; LacZ), was constructed in the same way.

All antigens were given in a soluble form, i.e. dissolved in either a HEPES buffer or in a buffer containing 0.5% Deoxycholate (DOC).

The antigens are listed in Table 5 below.

<u>Table 5</u> <u>Helicobacter pylori proteins</u>

Outer membrane Proteins

Protein 7116626 Protein 4721061 Protein 16225006 Protein 29479681 Protein 14640637

5 Periplasmic/Secreted Proteins

Protein 30100332

Other cell envelope proteins

Protein 4821082

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Flagella-associated proteins

Protein 26380318

Control proteins

15 b-galactosidase (LacZ)

Immunizations

Ten animals in each group were immunized 4 times over a 34 day period (day 1, 15, 25 and 35). Purified antigens in solution or suspension were given at a dose of 100 mg/mouse. As an adjuvant, the animals were also given 10 µg/mouse of Cholera toxin (CT) with each immunization. Omeprazole (400 mmol/kg) was given orally to the animals 3-5 h prior to immunization as a way of protecting the antigens from acid degradation. Infected control animals received HEPES buffer + CT or DOC buffer + CT. Animals were sacrificed 2-4 weeks after final immunization. A general outline of the study is shown in Table 6 below.

<u>Table 6</u> <u>Study outline, therapeutic immunization:</u>

Mice were all infected with H. pylori strain Ah244 at day 30.

	Substance	Mouse strain n=10	<u>Dose/mouse</u>	Dates for dosing
35	1. Controls, PBS	Balb/c	0.3 ml	0, 14, 24, 34
	2. Cholera toxin, 10 μg	Balb/c	0.3 ml	0, 14, 24, 34
40	3. Protein 16225006, 100 μg + CT 10	0 μg Balb/c	0.3 ml	0, 14, 24, 34
	4. Protein 26054702, 100 μg + CT 10	0 μg Balb/c	0.3 ml	0, 14, 24, 34

	5. Protein 26380318, 100 μg + CT 10 μg Balb/c	0.3 ml	0, 14, 24, 34
	6. Protein 29479681, 100 μg + CT 10 μg Balb/c	0.3 ml	0, 14, 24, 34
5	7. Protein 30100332, 100 μg + CT 10 μg Balb/c	0.3 ml	0, 14, 24, 34
	8. Protein 4721061, 100 μg + CT 10 μg Balb/c	0.3 ml	0, 14, 24, 34
	9. Protein 4821082, 100 μg + CT 10 μg Balb/c	0.3 ml	0, 14, 24, 34
10	10. Protein 7116626, 100 μg + CT 10 μg Balb/c	0.3 ml	0, 14, 24, 34
	11.Protein 14640637, 100 μg + CT 10 μg Balb/c	0.3 ml	0, 14, 24, 34

15 Analysis of infection

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Mucosal infection: The mice were sacrificed by CO₂ and cervical dislocation. The abdomen was opened and the stomach removed. After cutting the stomach along the greater curvature, it was rinsed in saline. The mucosa from the antrum and corpus of an area of 25mm² was scraped separately with a surgical scalpel. The mucosa scraping was suspended in Brucella broth and plated onto Blood Skirrow selective plates. The plates were incubated under microaerophilic conditions for 3-5 days and the number of colonies was counted. The identity of *H. pylori* was ascertained by urease and catalase test and by direct microscopy or Gram staining.

The urease test was performed essentially as follows. The reagent, Urea Agar Base Concentrate, was purchased from DIFCO Laboratories, Detroit, MI (Catalog # 0284-61-3). Urea agar base concentrate was diluted 1:10 with water. 1 ml of if the diluted concentrate was mixed with 100-200 ml of actively growing *H. pylori* cells. Color change to magenta indicated that cells were urease positive.

The catalase test was performed essentially as follows. The reagent, N,N,N',N'-Tetramethyl-p-Phenylenediamine, was purchased from Sigma, St. Louis, MO (Catalog # T3134). A solution of the regent (1% w/v in water) was prepared. *H. pylori* cells were swabbed onto Whatman filter paper and overlaid with the 1% solution. Color change to dark blue indicated that the cells were catalase positive.

<u>Serum antibodies:</u> From all mice serum was prepared from blood drawn by heart puncture. Serum antibodies were identified by regular ELISA techniques, where the specific antigens of *Helicobacter pylori* were plated.

Mucosal antibodies: Gentle scrapings of a defined part of the corpus and of 4 cm of duodenum were performed in 50% of the mice in order to detect the presence of

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antibodies in the mucous. The antibody titers were determined by regular ELISA technique as for serum antibodies.

Statistical analysis: Wilcoxon-Mann-Whitney sign rank test was used for determination of significant effects of the antigens on *Helicobacter pylori* colonization. P<0.05 was considered significant. Because the antrum is the major colonization site for *Helicobacter* most emphasis was put upon changes in the antral colonization.

Results

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Antibodies in sera: All antigens tested given together with CT gave rise to a measurable specific titer in serum. The highest responses were seen with Protein 7116626, Protein 4721061, Protein 26380318, Protein 14640637 and Protein 4821082 (see Figure 1).

Antibodies in mucus: In the mucus scrapings, specific antibodies against all antigens tested were seen. By far the strongest response was seen with Protein 30100332, followed by Protein 14640637, and Protein 26380318 (see Figure 2).

Therapeutic immunization effects:

All control animals (BALB/c mice) were well colonized with *H. pylori* (strain AH244) in both antrum and corpus of the stomach. Of the antigens tested 3 proteins (Protein 4721061, Protein 4821082, and Protein 14640637) gave a good and significant reduction and/or eradication of the *H. pylori* infection. The degree of colonization of the antrum was lower following immunization with Protein 7116626 and Protein 26380318 compared to control. The effect of Proteins 16225006, 29479681, and 30100332 did not differ from control. The control protein lacZ, i.e. the non-*H. pylori* protein, had no eradication effect and in fact had higher *Helicobacter* colonization compared to the HEPES + CT control. All data are shown in Figures 3 and 4 for proteins dissolved in HEPES and DOC respectively. Data is shown as geometric mean values. n=8-10 Wilcoxon-Mann-Whitney sign rank test * = p<0.05; x/10 = number of mice showing eradication of *H. pylori* over the total number of mice examined.

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The data presented indicate that all of the *H. pylori* associated proteins included in this study, when used as oral immunogens in conjunction with the oral adjuvant CT, resulted in stimulation of an immune response as measured by specific serum and mucosal antibodies. A majority of the proteins led to a reduction, and in some cases complete clearance of the colonization of *H. pylori* in this animal model. It should be noted that the reduction or clearance was due to heterologous protection rather than homologous protection (the polypeptides were based on the *H. pylori* J99 strain

sequence and used in the therapeutic immunization studies against a different (AH244) challenge strain, indicating the vaccine potential against a wide variety of *H. pylori* strains.

The highest colonization in the antrum was seen in animals treated with the non-Helicobacter protein LacZ, indicating that the effects seen with the Helicobacter pylori antigens were specific.

Taken together these data strongly support the use of these *H. pylori* proteins in a pharmaceutical formulation for the use in humans to treat and/or prevent *H. pylori* infections.

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V. Sequence Variance Analysis of genes in Helicobacter pylori strains

Four genes were cloned and sequenced from several strains of *H. pylori* to compare the DNA and deduced amino acid sequences. This information was used to determine the sequence variation between the *H. pylori* strain, J99, and other *H. pylori* strains isolated from human patients.

Preparation of Chromosomal DNA.

Cultures of *H. pylori* strains (as listed in Table 9) were grown in BLBB (1% Tryptone, 1% Peptamin 0.1% Glucose, 0.2% Yeast Extract 0.5% Sodium Chloride, 5% Fetal Bovine Serum) to an OD₆₀₀ of 0.2. Cells were centrifuged in a Sorvall RC-3B at 3500 x g at 4°C for 15 minutes and the pellet resuspended in 0.95 mls of 10 mM Tris-HCl, 0.1 mM EDTA (TE). Lysozyme was added to a final concentration of 1mg/ml along with, SDS to 1% and RNAse A + T1 to 0.5mg/ml and 5 units/ml respectively, and incubated at 37°C for one hour. Proteinase K was then added to a final concentration of 0.4mg/ml and the sample was incubated at 55 C for more than one hour. NaCl was added to the sample to a concentration of 0.65 M, mixed carefully, and 0.15 ml of 10% CTAB in 0.7M NaCL (final is 1% CTAB/70mM NaCL) was added followed by incubation at 65°C for 20 minutes. At this point, the samples were extracted with chloroform:isoamyl alcohol, extracted with phenol, and extracted again with chloroform:isoamyl alcohol. DNA was precipitated with either EtOH (1.5 x volumes) or isopropanol (0.6 x volumes) at -70°C for 10minutes, washed in 70% EtOH and resuspended in TE.

PCR Amplification and cloning.

Genomic DNA prepared from twelve strains of *Helicobacter pylori* was used as the source of template DNA for PCR amplification reactions (Current Protocols in Molecular Biology, John Wiley and Sons, Inc., F. Ausubel et al., editors, 1994). To

amplify a DNA sequence containing an *H. pylori* ORF, genomic DNA (10 nanograms) was introduced into a reaction vial containing 2 mM MgCl₂, 1 micromolar synthetic oligonucleotide primers (forward and reverse primers, see Table 7) complementary to and flanking a defined *H. pylori* ORF, 0.2 mM of each deoxynucleotide triphosphate; dATP, dGTP, dCTP, dTTP and 0.5 units of heat stable DNA polymerase (Amplitaq, Roche Molecular Systems, Inc., Branchburg, NJ, USA) in a final volume of 20 microliters in duplicate reactions.

Table 7

Oligonucleotide primers used for PCR amplification of *H. pylori* DNA sequences.

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Outer membrane	Forward primer 5' to 3'	Reverse Primer 5' to 3'
	Forward primer 5 to 5	Reverse Filmer 5 to 5
Proteins		
D : : 2/05/1502/5	5'-	5'-
Protein 26054702 (for	1 ~	1 -
strains AH4, AH15,	TTAACCATGGTGAAA	TAGAATTCGCCTCTA
AH61, 5294, 5640,	AGCGATA-3' (SEQ ID	AAACTTTAG-3' (SEQ
AH18, and AH244)	NO:169)	ID NO:170)
Protein 26054702	5'-	5'-
(for strains AH5, 5155,	TTAACCATGGTGAAA	TAGAATTCGCATAAC
7958, AH24,and J99)	AGCGATA-3' (SEQ ID	GATCAATC-3' (SEQ ID
	NO:171)	NO:172)
-		
Protein 7116626	5'-	5'-
	ATATCCATGGTGAGT	ATGAATTCAATTTTT
	TTGATGA-3' (SEQ ID	TATTTTGCCA-3' (SEQ
	NO:173)	ID NO:174)
Protein 29479681	5'-	5'-
	AATTCCATGGCTATC	ATGAATTCGCCAAAA
	CAAATCCG-3' (SEQ ID	TCGTAGTATT-3' (SEQ
	NO:175)	ID NO:176)
		12 1.0.1.0)
Protein 346	5'-	5'-
	GATACCATGGAATTT	TGAATTCGAAAAAGT
	ATGAAAAAG-3' (SEQ	GTAGTTATAC-3' (SEQ
	ID NO:177)	ID NO:178)
	וויטאו עו	(פו זיחעו חד ו

The following thermal cycling conditions were used to obtain amplified DNA products for each ORF using a Perkin Elmer Cetus/ GeneAmp PCR System 9600 thermal cycler:

Protein 7116626 and Protein 346;
Denaturation at 94°C for 2 min,
2 cycles at 94°C for 15 sec, 30°C for 15 sec and 72°C for 1.5 min
5 23 cycles at 94°C for 15 sec, 55°C for 15 sec and 72°C for 1.5 min
Reactions were concluded at 72°C for 6 minutes.

Protein 26054702 for strains AH5, 5155, 7958, AH24, and J99; Denaturation at 94°C for 2 min,

2 cycles at 94°C for 15 sec, 30°C for 15 sec and 72°C for 1.5 min 25 cycles at 94°C for 15 sec, 55°C for 15 sec and 72°C for 1.5 min Reaction was concluded at 72°C for 6 minutes.

Protein 26054702 and Protein 294796813 for strains AH4, AH15, AH61, 5294, 5640,

AH18, and Hp244;

Denaturation at 94°C for 2 min, 2 cycles at 94°C for 15 sec, 30°C for 20 sec and 72°C for 2 min 25 cycles at 94°C for 15 sec, 55°C for 20 sec and 72°C for 2 min Reactions were concluded at 72°C for 8 minutes.

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Upon completion of thermal cycling reactions, each pair of samples were combined and used directly for cloning into the pCR cloning vector as described below.

Cloning of H. pylori DNA sequences into the pCR TA cloning vector.

All amplified inserts were cloned into the pCR 2.1 vector by the method described in the Original TA cloning kit (Invitrogen, San Diego, CA). Products of the ligation reaction were then used to transform the TOP10F' (INVaF' in the case of *H. pylori* sequence 350) strain of *E. coli* as described below.

Transformation of competent bacteria with recombinant plasmids

Competent bacteria, *E coli* strain TOP10F' or *E. coli* strain INVaF' were transformed with recombinant pCR expression plasmids carrying the cloned *H. pylori* sequences according to standard methods (Current Protocols in Molecular Biology, John Wiley and Sons, Inc., F. Ausubel et al., editors, 1994). Briefly, 2 microliters of 0.5 micromolar BME was added to each vial of 50 microliters of competent cells. Subsequently, 2 microliters of ligation reaction was mixed with the competent cells and incubated on ice for 30 minutes. The cells and ligation mixture were then subjected to a

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"heat shock" at 42°C for 30 seconds, and were subsequently placed on ice for an additional 2 minutes, after which, samples were incubated in 0.45 milliliters SOC medium (0.5% yeast extract, 2.0 % tryptone, 10 mM NaCl, 2.5 mM KCl, 10 mM MgCl2, 10 mM MgSO4 and 20, mM glucose) at 37°C with shaking for 1 hour. Samples were then spread on LB agar plates containing 25 microgram/ml kanamycin sulfate or 100 micrograms/ml ampicillan for growth overnight. Transformed colonies of TOP10F' or INVaF' were then picked and analyzed to evaluate cloned inserts as described below. *Identification of recombinant PCR plasmids carrying H. pylori sequences*

Individual TOP10F' or INVaF' clones transformed with recombinant pCR-H.pylori ORFs were analyzed by PCR amplification of the cloned inserts using the same forward and reverse primers, specific for each H. pylori sequence, that were used in the original PCR amplification cloning reactions. Successful amplification verified the integration of the H. pylori sequences in the cloning vector (Current Protocols in Molecular Biology, John Wiley and Sons, Inc., F. Ausubel et al., editors, 1994).

Individual clones of recombinant pCR vectors carrying properly cloned *H. pylori* ORFs were picked for sequence analysis. Sequence analysis was performed on ABI Sequencers using standard protocols (Perkin Elmer) using vector-specific primers (as found in PCRII or pCR2.1, Invitrogen, San Diego, CA) and sequencing primers specific to the ORF as listed in Table 8 below.

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<u>Table 8</u>
<u>Oligonucleotide primers used for sequencing of *H. pylori* DNA sequences.</u>

Outer membrane	Forward primers 5' to 3'	Reverse Primers 5' to 3'
Proteins		
Protein 26054702	5'-	5'-
	CCCTTCATTTTAGAAATC G-3' (SEQ ID NO:179)	CTTTGGGTAAAAACGCA TC-3' (SEQ ID NO:186)
	5'-	5'-
	ATTTCAACCAATTCAAT GCG-3' (SEQ ID NO:180) 5'-	CGATCTTTGATCCTAATT CA-3' (SEQ ID NO:187) 5'-
	GCCCCTTTTGATTTGAAG CT-3' (SEQ ID NO:181) 5'-	ATCAAGTTGCCTATGCT GA-3' (SEQ ID NO:188)
	TCGCTCCAAGATACCAA GAAGT-3' (SEQ ID NO:182) 5'-	
	CTTGAATTAGGGGCAAA GATCG-3' (SEQ ID NO:183)	
	5'- ATGCGTTTTTACCCAAA GAAGT-3' (SEQ ID NO:184) 5'-	
	ATAACGCCACTTCCTTAT TGGT-3' (SEQ ID NO:185)	
D	61	
Protein 7116626	5'- TTGAACACTTTTGATTAT GCGG-3' (SEQ ID NO:189) 5'-	5'- GTCTTTAGCAAAAATGG CGTC-3' (SEQ ID NO:191) 5'-
	GGATTATGCGATTGTTTT ACAAG-3' (SEQ ID NO:190)	AATGAGCGTAAGAGAGC CTTC-3' (SEQ ID NO:192)
Protein	5'-	5'-
29479681	CTTATGGGGGTATTGTC A-3' (SEQ ID NO:193) 5'-	AGGTTGTTGCCTAAAGA CT-3' (SEQ ID NO:195) 5'-
	AGCATGTGGGTATCCAG C-3' (SEQ ID NO:194)	CTGCCTCCACCTTTGATC -3' (SEQ ID NO:196)

Protein 346	5'- ACCAATATCAATTGGCA CT-3' (SEQ ID NO:197) 5'- ACTTGGAAAAGCTCTGC A-3' (SEQ ID NO:198)	5'- CTTGCTTGTCATATCTAG C-3' (SEQ ID NO:199) 5'- GTTGAAGTGTTGGTGCT A-3' (SEQ ID NO:200)
	5'- CAAGCAAGTGGTTTGGT TTTAG-3' (SEQ ID NO:201) 5'- TGGAAAGAGCAAATCAT TGAAG-3' (SEQ ID NO:202)	5'- GCCCATAATCAAAAAGC CCAT-3' (SEQ ID NO:203) 5'- CTAAAAACCAAACCACTT GCT TGTC-3' (SEQ ID NO:204)
Vector Primers	5'- GTAAAACGACGGCCAG- 3' (SEQ ID NO:205)	5'- CAGGAAACAGCTATGAC -3' (SEQ ID NO:206)

Results

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To establish the PCR error rate in these experiments, five individual clones of Protein 26054702, prepared from five separate PCR reaction mixtures from *H. pylori* strain J99, were sequenced over a total length of 897 nucleotides for a cumulative total of 4485 bases of DNA sequence. DNA sequence for the five clones was compared to a DNA sequence obtained previously by a different method, i.e., random shotgun cloning and sequencing. The PCR error rate for the experiments described herein was determined to be 2 base changes out of 4485 bases, which is equivalent to an estimated error rate of less than or equal to 0.04%.

DNA sequence analysis was performed on four different open reading frames identified as genes and amplified by PCR methods from a dozen different strains of the bacterium *Helicobacter pylori*. The deduced amino acid sequences of three of the four open reading frames that were selected for this study showed statistically significant BLAST homology to defined proteins present in other bacterial species. Those ORFs included: Protein 26054702, homologous to the val A & B genes encoding an ABC transporter in F. novicida; Protein 7116626, homologous to lipoprotein e (P4) present in the outer membrane of H. influenzae; Protein 29479681, homologous to fecA, an outer membrane receptor in iron (III) dicitrate transport in E. coli. Protein 346 was identified as an unknown open reading frame, because it showed low homology with sequences in the public databases.

To assess the extent of conservation or variance in the ORFs across various strains of *H. pylori*, changes in DNA sequence and the deduced protein sequence were compared to the DNA and deduced protein sequences found in the J99 strain of *H.*

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pylori (see Table 9 below). Results are presented as percent identity to the J99 strain of H. pylori sequenced by random shotgun cloning. To control for any variations in the J99 sequence each of the four open reading frames were cloned and sequenced again from the J99 bacterial strain and that sequence information was compared to the sequence information that had been collected from inserts cloned by random shotgun sequencing of the J99 strain. The data demonstrate that there is variation in the DNA sequence ranging from as little as 0.12 % difference (Protein 346, J99 strain) to approximately 7% change (Protein 26054702, strain AH5). The deduced protein sequences show either no variation (Protein 346, strains AH18 and AH24) or up to as much as 7.66% amino acid changes (Protein 26054702, Strain AH5).

Table_9

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Multiple Strain DNA Sequence analysis of H. pylori Vaccine Candidates								
J99 Protein #:	26054702	2054702	7116626 7	116626	29479681	29479681	346 3	46
Length of Reg Sequenced:	Length of Region Sequenced: 248 a.a. 746 nt. 232 a.a. 96 nt. 182 a.a. 548 nt. 273 a.a. 819 nt.							9 nt.
Strain Tested	AA identity	Nuc.	AA identity	Nuc.	AA identity	Nuc. identity	AA identity	Nuc.
J99	100.00%	100.00%	100.00%	100.00%	6 100.00%	100.00%	99.63%	99.88%
AH244	95.16%	95.04%	n.d.	n.d.	99.09%	96.71%	98.90%	96.45%
AH4	95.97%	95.98%	97.84%	95.83%	n.d.	n.d.	97.80%	95.73%
AH5	92.34%	93.03%	98.28%	96.12%	98.91%	96.90%	98.53%	95.73%
AH15	95.16%	94.91%	97.41%	95.98%	99.82%	9 7.99 %	99.63%	96.09%
AH61	n.d.	n.d.	97.84%	95.98%	99.27%	97.44%	n.d.	n.d.
5155	n.d.	n.d.	n.d.	n.d.	99.45%	97.08%	98.53%	95.60%
5294	94.35%	94.37%	98.28%	95.40%	99.64%	97.26%	97.07%	95.48%
7958	94.35%	94.10%	97.84%	95.40%	n.d.	n.d.	99.63%	96.46%
5640	95.16%	94.37%	97.41%	95.69%	99.09%	97.63%	98.53%	95.48%
AH18	n.d.	n.d.	98.71%	95.69%	99.64%	97.44%	100.00%	95.97%
AH24	94.75%	95.04%	97.84%	95.40%	99.27%	96.71%	100.00%	96.46%

n.d.= not done.

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VI. Experimental Knock-Out Protocol for the Determination of Essential H. pylori Genes as Potential Therapeutic Targets

Therapeutic targets are chosen from genes whose protein products appear to play key roles in essential cell pathways such as cell envelope synthesis, DNA synthesis, transcription, translation, regulation and colonization/virulence.

The protocol for the deletion of portions of *H. pylori* genes/ORFs and the insertional mutagenesis of a kanamycin-resistance cassette in order to identify genes which are essential to the cell is modified from previously published methods (Labigne-Roussel et al., 1988, J. Bacteriology 170, pp. 1704-1708; Cover et al.,1994, J. Biological Chemistry 269, pp. 10566-10573; Reyrat et al., 1995, Proc. Natl. Acad. Sci. 92, pp 8768-8772). The result is a gene "knock-out."

Identification and Cloning of H. pylori Gene Sequences

The sequences of the genes or ORFs (open reading frames) selected as knock-out targets are identified from the *H. pylori* genomic sequence and used to design primers to specifically amplify the genes/ORFs. All synthetic oligonucleotide primers are designed with the aid of the OLIGO program (National Biosciences, Inc., Plymouth, MN 55447, USA), and can be purchased from Gibco/BRL Life Technologies (Gaithersburg, MD, USA). If the ORF is smaller than 800 to 1000 base pairs, flanking primers are chosen outside of the open reading frame.

Genomic DNA prepared from the *Helicobacter pylori* HpJ99 strain (ATCC 55679; deposited by Genome Therapeutics Corporation, 100 Beaver Street, Waltham, MA 02154) is used as the source of template DNA for amplification of the ORFs by PCR (polymerase chain reaction) (Current Protocols in Molecular Biology, John Wiley and Sons, Inc., F. Ausubel et al., editors, 1994). For the preparation of genomic DNA from *H. pylori*, see Example I. PCR amplification is carried out by introducing 10 nanograms of genomic HpJ99 DNA into a reaction vial containing 10 mM Tris pH 8.3, 50 mM KCl, 2 mM MgCl₂, 2 microMolar synthetic oligonucleotide primers (forward=F1 and reverse=R1), 0.2 mM of each deoxynucleotide triphosphate (dATP,dGTP, dCTP, dTTP), and 1.25 units of heat stable DNA polymerase (Amplitaq, Roche Molecular Systems, Inc., Branchburg, NJ, USA) in a final volume of 40 microliters. The PCR is carried out with Perkin Elmer Cetus/GeneAmp PCR System 9600 thermal cyclers.

Upon completion of thermal cycling reactions, each sample of amplified DNA is visualized on a 2% TAE agarose gel stained with Ethidium Bromide (Current Protocols in Molecular Biology, John Wiley and Sons, Inc., F. Ausubel et al., editors, 1994) to

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determine that a single product of the expected size had resulted from the reaction. Amplified DNA is then washed and purified using the Qiaquick Spin PCR purification kit (Qiagen, Gaithersburg, MD, USA).

PCR products are cloned into the pT7Blue T-Vector (catalog#69820-1, Novagen, Inc., Madison, WI, USA) using the TA cloning strategy (Current Protocols in Molecular Biology, John Wiley and Sons, Inc., F. Ausubel et al., editors, 1994). The ligation of the PCR product into the vector is accomplished by mixing a 6 fold molar excess of the PCR product, 10 ng of pT7Blue-T vector (Novagen), 1 microliter of T4 DNA Ligase Buffer (New England Biolabs, Beverly, MA, USA), and 200 units of T4 DNA Ligase (New England Biolabs) into a final reaction volume of 10 microliters. Ligation is allowed to proceed for 16 hours at 16°C.

Ligation products are electroporated (Current Protocols in Molecular Biology, John Wiley and Sons, Inc., F. Ausubel et al., editors, 1994) into electroporation-competent XL-1 Blue or DH5-a *E.coli* cells (Clontech Lab., Inc. Palo Alto, CA, USA). Briefly, 1 microliter of ligation reaction is mixed with 40 microliters of electrocompetent cells and subjected to a high voltage pulse (25 microFarads, 2.5 kV, 200 ohms) after which the samples are incubated in 0.45 ml SOC medium (0.5% yeast extract, 2% tryptone, 10 mM NaCl, 2.5 mM KCl, 10 mM MgCl₂, 10 mM MgSO₄ and 20 mM glucose) at 37°C with shaking for 1 hour. Samples are then spread onto LB (10 g/l bacto tryptone, 5 g/l bacto yeast extract, 10 g/l sodium chloride) plates containing 100 microgram/ml of Ampicillin, 0.3% X-gal, and 100 microgram/ml IPTG. These plates are incubated overnight at 37°C. Ampicillin-resistant colonies with white color are selected, grown in 5 ml of liquid LB containing 100 microgram/ml of Ampicillin, and plasmid DNA is isolated using the Qiagen miniprep protocol (Qiagen, Gaithersburg, MD, USA).

To verify that the correct *H.pylori* DNA inserts had been cloned, these pT7Blue plasmid DNAs are used as templates for PCR amplification of the cloned inserts, using the same forward and reverse primers used for the initial amplification of the J99 *H.pylori* sequence. Recognition of the primers and a PCR product of the correct size as visualized on a 2% TAE, ethidium bromide stained agarose gel are confirmation that the correct inserts had been cloned. Two to six such verified clones are obtained for each knock-out target, and frozen at -70°C for storage. To minimize errors due to PCR, plasmid DNA from these verified clones are pooled, and used in subsequent cloning steps.

The sequences of the genes/ORFs are again used to design a second pair of primers which flank the region of *H. pylori* DNA to be either interrupted or deleted (up to 250 basepairs) within the ORFs but are oriented away from each other. The pool of

circular plasmid DNAs of the previously isolated clones are used as templates for this round of PCR. Since the orientation of amplification of this pair of deletion primers is away from each other, the portion of the ORF between the primers is not included in the resultant PCR product. The PCR product is a linear piece of DNA with *H. pylori* DNA at each end and the pT7Blue vector backbone between them which, in essence, resultes in the deletion of a portion of the ORFs. The PCR product is visualized on a 1% TAE, ethidium bromide stained agarose gel to confirm that only a single product of the correct size has been amplified.

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A Kanamycin-resistance cassette (Labigne-Roussel et al., 1988 J. Bacteriology 170, 1704-1708) is ligated to this PCR product by the TA cloning method used previously (Current Protocols in Molecular Biology, John Wiley and Sons, Inc., F. Ausubel et al., editors, 1994). The Kanamycin cassette containing a Campylobacter kanamycin resistance gene is obtained by carrying out an EcoRI digestion of the recombinant plasmid pCTB8:kan (Cover et al., 1994, J. Biological Chemistry 269, pp. 10566-10573). The proper fragment (1.4 kb) is isolated on a 1% TAE gel, and isolated using the QIAquick gel extraction kit (Qiagen, Gaithersburg, MD, USA). The fragment is end repaired using the Klenow fill-in protocol, which involved mixing 4ug of the DNA fragment, 1 microliter of dATP, dCTP, dCTP, dTTP at 0.5 mM, 2 microliter of Klenow Buffer (New England Biolabs) and 5 units of Klenow DNA Polymerase I Large (Klenow) Fragment (New England Biolabs) into a 20 microliter reaction, incubating at 30°C for 15 min, and inactivating the enzyme by heating to 75°C for 10 minutes. This blunt-ended Kanamycin cassette is then purified through a Qiaquick column (Qiagen, Gaithersburg, MD, USA) to eliminate nucleotides. The "T" overhang is then generated by mixing 5 micrograms of the blunt-ended kanamycin cassette, 10 mM Tris pH 8.3, 50 mM KCl, 2 mM MgCl₂, 5 units of DNA Polymerase (Amplitaq, Roche Molecular Systems, Inc., Branchburg, NJ, USA), 20 microliters of 5 mM dTTP, in a 100 microliter reaction and incubating the reaction for 2 hours at 37°C. The "Kan-T" cassette is purified using a QIAquick column (Qiagen, Gaithersburg, MD, USA). The PCR product of the deletion primers (F2 and R2) is ligated to the Kan-T cassette by mixing 10 to 25 ng of deletion primer PCR product, 50 - 75 ng Kan-T cassette DNA, 1 microliter 10x T4 DNA Ligase reaction mixture, 0.5 microliter T4 DNA Ligase (New England Biolabs, Beverly, MA, USA) in a 10 microliter reaction and incubating for 16 hours at 16°C.

The ligation products are transformed into XL-1 Blue or DH5-a *E.coli* cells by electroporation as described previously. After recovery in SOC, cells are plated onto LB plates containing 100 microgram/ml Ampicillin and grown overnight at 37°C. These plates are then replica plated onto plates containing 25 microgram/ml Kanamycin and

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allowed to grow overnight. Resultant colonies have both the Ampicillin resistance gene present in the pT7Blue vector, and the newly introduced Kanamycin resistance gene. Colonies are picked into LB containing 25 microgram/ml Kanamycin and plasmid DNA is isolated from the cultured cells using the Qiagen miniprep protocol (Qiagen, Gaithersburg, MD, USA).

Several tests by PCR amplification are conducted on these plasmids to verify that the Kanamycin is inserted in the H. pylori gene/ORF, and to determine the orientation of the insertion of the Kanamycin-resistance gene relative to the H. pylori gene/ORF. To verify that the Kanamycin cassette is inserted into the H. pylori sequence, the plasmid DNAs are used as templates for PCR amplification with the set of primers originally used to clone the H. pylori gene/ORFs. The correct PCR product is the size of the deleted gene/ORF but increased in size by the addition of a 1.4 kilobase Kanamycin cassette. To avoid potential polar effects of the kanamycin resistance cassette on H. pylori gene expression, the orientation of the Kanamycin resistance gene with respect to the knock-out gene/ORF is determined and both orientations are eventually used in H. pylori transformations (see below). To determine the orientation of insertion of the kanamycin resistance gene, primers are designed from the ends of the kanamycin resistance gene ("Kan-1" 5'-ATCTTACCTATCACCTCAAAT-3' (SEQ ID NO:207)), and "Kan-2" 5'-AGACAGCAACATCTTTGTGAA-3' (SEQ ID NO:208)). By using each of the cloning primers in conjunction with each of the Kan primers (4 combinations of primers), the orientation of the Kanamycin cassette relative to the H.pylori sequence is determined. Positive clones are classified as either in the "A" orientation (the same direction of transcription is present for both the H. pylori gene and the Kanamycin resistance gene), or in the "B" orientation (the direction of transcription for the H.pylori gene is opposite to that of the Kanamycin resistance gene). Clones which share the same orientation (A or B) are pooled for subsequent experiments and independently transformed into H. pylori.

Transformation of Plasmid DNA into H. pylori cells

Two strains of *H. pylori* are used for transformation: ATCC <u>55679</u>, the clinical isolate which provided the DNA from which the *H. pylori* sequence database is obtained, and AH244, an isolate which had been passaged in, and has the ability to colonize the mouse stomach. Cells for transformation are grown at 37°C, 10% CO₂, 100% humidity, either on Sheep-Blood agar plates or in Brucella Broth liquid. Cells are grown to exponential phase, and examined microscopically to determine that the cells are "healthy" (actively moving cells) and not contaminated. If grown on plates, cells are harvested by scraping cells from the plate with a sterile loop, suspended in 1 ml of

Brucella Broth, spun down (1 minute, top speed in eppendorf microfuge) and resuspended in 200 microliters Brucella Broth. If grown in Brucella Broth liquid, cells are centrifuged (15 minutes at 3000 rpm in a Beckman TJ6 centrifuge) and the cell pellet resuspended in 200 microliters of Brucella broth. An aliquot of cells is taken to determine the optical density at 600 nm, in order to calculate the concentration of cells. An aliquot (1 to 5 OD₆₀₀ units/25 microliter) of the resuspended cells is placed onto a prewarmed Sheep-Blood agar plate, and the plate is further incubated at 37°C, 6% CO₂, 100% humidity for 4 hours. After this incubation, 10 microliters of plasmid DNA (100 micrograms per microliter) is spotted onto these cells. A positive control (plasmid DNA with the ribonuclease H gene disrupted by kanamycin resistance gene) and a negative control (no plasmid DNA) are done in parallel. The plates are returned to 37°C, 6% CO₂ for an additional 4 hours of incubation. Cells are then spread onto that plate using a swab wetted in Brucella broth, and grown for 20 hours at 37°C, 6% CO₂. Cells are then transferred to a Sheep-Blood agar plate containing 25 micrograms/ml Kanamycin, and allowed to grow for 3 to 5 days at 37°C, 6% CO₂, 100% humidity. If colonies appear, they are picked and regrown as patches on a fresh Sheep-Blood agar plate containing 25 micrograms/ml Kanamycin.

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Three sets of PCR tests are done to verify that the colonies of transformants have arisen from homologous recombination at the proper chromosomal location. The template for PCR (DNA from the colony) is obtained by a rapid boiling DNA preparation method as follows. An aliquot of the colony (stab of the colony with a toothpick) is introduced into 100 microliters of 1% Triton X-100, 20 mM Tris, pH 8.5, and boiled for 6 minutes. An equal volume of phenol: chloroform (1:1) is added and vortexed. The mixture is microfuged for 5 minutes and the supernatant is used as DNA template for PCR with combinations of the following primers to verify homologous recombination at the proper chromosomal location.

TEST 1. PCR with cloning primers originally used to amplify the gene/ORF. A positive result of homologous recombination at the correct chromosomal location should show a single PCR product whose size is expected to be the size of the deleted gene/ORF but increased in size by the addition of a 1.4 kilobase Kanamycin cassette. A PCR product of just the size of the gene/ORF is proof that the gene had not been knocked out and that the transformant is not the result of homologous recombination at the correct chromosome location.

TEST 2. PCR with F3 (primer designed from sequences upstream of the gene/ORF and not present on the plasmid), and either primer Kan-1 or Kan-2 (primers designed from the ends of the kanamycin resistance gene), depending on whether the plasmid DNA used was of "A" or "B" orientation. Homologous recombination at the

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correct chromosomal location will result in a single PCR product of the expected size (i.e., from the location of F3 to the insertion site of kanamycin resistance gene). No PCR product or PCR product(s) of incorrect size(s) will prove that the plasmid had not integrated at the correct site and that the gene had not been knocked out.

TEST 3. PCR with R3 (primer designed from sequences downstream of the gene/ORF and not present on the plasmid) and either primer Kan-1 or Kan-2, depending on whether the plasmid DNA used was of "A" or "B" orientation. Homologous recombination at the correct chromosomal location will result in a single PCR product of the expected size (i.e., from the insertion site of kanamycin resistance gene to the downstream location of R3). Again, no PCR product or PCR product(s) of incorrect size(s) will prove that the plasmid had not integrated at the correct site and that the gene had not been knocked out.

Transformants showing positive results for all three tests above indicate that the gene is not essential for survival *in vitro*.

A negative result in any of the three above tests for each transformant indicates that the gene had not been disrupted, and that the gene is essential for survival in vitro.

In the event that no colonies result from two independent transformations while the positive control with the disrupted ribonuclease H plasmid DNA produces transformants, the plasmid DNA is further analyzed by PCR on DNA from transformant populations prior to plating for colony formation. This will verify that the plasmid can enter the cells and undergo homologous recombination at the correct site. Briefly, plasmid DNA is incubated according to the transformation protocol described above. DNA is extracted from the *H. pylori* cells immediately after incubation with the plasmid DNAs and the DNA is used as template for the above TEST 2 and TEST 3. Positive results in TEST 2 and TEST 3 would verify that the plasmid DNA could enter the cells and undergo homologous recombination at the correct chromosomal location. If TEST 2 and TEST 3 are positive, then failure to obtain viable transformants indicates that the gene is essential, and cells suffering a disruption in that gene are incapable of colony formation

VII. High-throughput drug screen assay

Cloning, expression and protein purification

Cloning, transformation, expression and purification of the *H. pylori* target gene and its protein product, e.g., an *H. pylori* enzyme, to be used in a high-throughput drug screen assay, is carried out essentially as described in Examples II and III above. Development and application of a screening assay for a particular *H. pylori* gene product, peptidyl-propyl *cis-trans* isomerase, is described below as a specific example.

Enzymatic Assay

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The assay is essentially as described by Fisher (Fischer, G., et.al. (1984) *Biomed. Biochim. Acta* 43:1101-1111). The assay measures the *cis-trans* isomerization of the Ala-Pro bond in the test peptide N-succinyl-Ala-Ala-Pro-Phe-p-nitroanilide (Sigma # S-7388, lot # 84H5805). The assay is coupled with α-chymotrypsin, where the ability of the protease to cleave the test peptide occurs only when the Ala-Pro bond is in *trans*. The conversion of the test peptide to the trans isomer in the assay is followed at 390 nm on a Beckman Model DU-650 spectophotometer. The data are collected every second with an average scanning of time of 0.5 second. Assays are carried out in 35 mM Hepes, pH 8.0, in a final volume of 400 ul, with 10 μM α-chymotrypsin (type 1-5 from bovine-Pancreas, Sigma #-C=7762, lot-23H7020) and 10 nM-PPIase. To initiate the reaction, 10 μl of the substrate (2 mM N-Succinyl-Ala-Ala-Pro-Phe-p-nitroanilide in DMSO) is added to 390 μl of reaction mixture at room temperature.

15 Enzymatic assay in crude bacterial extract.

A 50 ml culture of *Helicobacter pylori* (strain J99) in Brucella broth is harvested at mid-log phase (OD $_{600~nm} \sim 1$) and resuspended in lysis buffer with the following protease inhibitors: 1 mM PMSF, and 10 µg/ml of each of aprotinin, leupeptin, pepstatine, TLCK, TPCK, and soybean trypsin inhibitor. The suspension is subjected to 3 cycles of freeze-thaw (15 minutes at -70 $^{\circ}$ C, then 30 minutes at room temperature), followed by sonication (three 20 second bursts). The lysate is centrifuged (12,000 g x 30 minutes) and the supernatant is assayed for enzymatic activity as described above.

Many H. pylori enzymes can be expressed at high levels and in an active form in E. coli. Such high yields of purified proteins provide for the design of various high throughput drug screening assays.

EQUIVALENTS

Those skilled in the art will recognize, or be able to ascertain using no more than routine experimentation, many equivalents to the specific embodiments and methods described herein. Such equivalents are intended to be encompassed by the scope of the following claims.

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SEQUENCE LISTING

5	1) GENERAL INFORMATION:
	(i) APPLICANT:
	(A) NAME: Astra Aktiebolag (B) STREET: S-151 85
	(C) CITY: Sodertalje
10	(D) STATE:
	(E) COUNTRY: Sweden (F) POSTAL CODE (ZIP)
	(F) POSTAL CODE (ZIP)
	(ii) TITLE OF INVENTION: NUCLEIC ACID AND AMINO ACID SEQUENCES
15	RELATING TO HELICOBACTER PYLORI AND
	VACCINE COMPOSITIONS THEREOF
20	(iii) NUMBER OF SEQUENCES: 208
20	(iv) COMPUTER READABLE FORM:
	(A) MEDIUM TYPE:
	(B) COMPUTER: (C) OPERATING SYSTEM:
25	(D) SOFTWARE:
23	
	(v) CURRENT APPLICATION DATA:
	(A) APPLICATION NUMBER (B) FILING DATE:
30	(B) FILING DATE:
30	(vi) PRIOR APPLICATION DATA:
	(A) APPLICATION NUMBER:US 08/739,150
	(B) FILING DATE: 28-OCT-1996
35	(vii) PRIOR APPLICATION DATA:
	(A) APPLICATION NUMBER: US 08/759,739
	(B) FILING DATE: 06-DEC-1996
	(viii) PRIOR APPLICATION DATA:
40	(A) APPLICATION NUMBER: US 08/891,928
	(B) FILING DATE: 14-JULY-1997
	(ix) CORRESPONDENCE ADDRESS:
	(A) ADDRESSEE: LAHIVE & COCKFIELD
45	(B) STREET: 28 State Street
	(C) CITY: Boston (D) STATE: Massachusetts
	(E) COUNTRY: USA
	(F) ZIP: 02109-1875
50	
	(x) ATTORNEY/AGENT INFORMATION:
	(A) NAME: Mandragouras, Amy E.(B) REGISTRATION NUMBER: 36,207
	(C) REFERENCE/DOCKET NUMBER: GTN-001CP10PC
55	(0,)====================================

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	(xi) TELECOMMUNICATION INFORMATION: (A) TELEPHONE: (617)227-7400 (B) TELEFAX: (617)742-4214	,
5	(2) INFORMATION FOR SEQ ID NO:1:	
10	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 561 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
	(ii) MOLECULE TYPE: DNA (genomic)	
15	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
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50	(iv) ANTI-SENSE: NO	
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	(C) STRANDEDNESS: double	
20	(D) TOPOLOGY: circular	
	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
25	(
	(iv) ANTI-SENSE: NO	
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(2) INFORMATION FOR SEQ ID NO:4:

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(i) SEQUENCE CHARACTERISTICS:
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               (C) STRANDEDNESS: double
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         (ii) MOLECULE TYPE: DNA (genomic)
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        (iii) HYPOTHETICAL: NO
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        (vi) ORIGINAL SOURCE:
               (A) ORGANISM: Helicobacter pylori
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               (B) LOCATION 1...831
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               (D) TOPOLOGY: circular
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        (iv) ANTI-SENSE: NO
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25	(C) STRANDEDNESS: double	
2.5	(D) TOPOLOGY: circular	
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	GATTATAAAG AGCAATTAGG CATGCTCGCA ACTTTCATTA GCCCTAATTC GCCCGTGATT	720
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1 1	MARIALITATE ATOMICOCCI COMMINGUILL TOTAL COMMINGUILL	

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20	(ii) MOLECULE TYPE: DNA (genomic) (iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
25	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
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55	(iv) ANTI-SENSE: NO	

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30	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1179 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
35	(ii) MOLECULE TYPE: DNA (genomic) (iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
40	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
45	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 11179</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:10:	
50	ATGAGAAAAC TATTCATCCC ACTTTTATTA TTCAGCGCTT TAGAAGCGAA CGAGAAAAAC GGCTTTTTCA TAGAAGCCGG CTTTGAAACT GGGCTATTAG AAGGCACACA AACGCAAGAA AAAAGACACA CCACCACAAA AAACACTTAC GCAACTTACA ATTATTTACC CACAGACACG ATTTTAAAAA GAGCGGCTAA TTTATTCACC AATGCCGAAG CGATTTCAAA ATTAAAATTC TCATCTTTAT CCCCTGTTAG AGTGTTGTAT ATGTATAATG GTCAATTAAC TATAGAAAAC	120 180 240 300
55	TTCTTGCCTT ATAATTTAAA TAATGTTAAG CTTAGTTTTA CAGACGCTCA AGGCAATGTG ATCGATCTAG GCGTGATAGA GACTATCCCC AAACACTCTA AGATTGTTTT GCCCGGAGAG	360 420

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	GCATTTGATA GTCTAAAAAT TGACCCCTAT ACTTTATTTC TTCCAAAAAT TGAAGCCACT	480
	AGCACTTCTA TTTCTGACGC TAACACGCAG AGGGTGTTTG AAACGCTCAA TAAGATTAAC	540
	ACAAATTTGG TCGTAAATTA TAGGAATGAA AACAAATTTA AAGATCACGA AAATCATTGO	600
	GAAGCCTTTA CCCCACAAAC CGCAGAAGAA TTCACTAATT TAATGTTGAA CATGATCGCT	660
5	GAAGCCITTA CCCAATCTTG GGGCGATGCG ATCTTAAACG CTCCTTTTGA GTTCACTAAC	720
	AGCCCAACAG ATTGCGATAA TGATCCTTCA AAATGCGTAA ATCCTGGGAC AAACGGGCTT	780
	GTCAATTCTA AAGTCGATCA AAAATATGTG TTAAACAAAC AAGACATTGT CAATAAATT	840
	AAAAACAAAG CGGATCTTGA TGTAATTGTT TTAAAGGATT CAGGGGTTGT AGGGCTTGGC	900
	AGAGATATTA CCCCTAGCAA CAATGATGAT GGCAAGCATT ATGGCCAGTT AGGGGTAGTA	960
10	GCTTCTGCTT TAGATCCTAA AAAACTCTTT GGCGATAACC TTAAGACTAT CAATTTAGAC	1020
10	GATTTAAGAA CCATCTTGCA TGAATTCAGC CACACTAAAG GCTATGGGCA TAACGGGAA	1080
	ATGACCTATC AAAGAGTGCC GGTAACGAAA GATGGTCAAG TGGAAAAGGA TAGTAATGG	1140
	AAGCCAAAAG ATTCTGATGG CCTCCCCTAT AATGTGTGT	1179
	AAGCCAAAAG ATTCTGATGG CCTCCCCTTT TITTCTCT	
15	(2) INFORMATION FOR SEQ ID NO:11:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 813 base pairs	
	(B) TYPE: nucleic acid	
20	(C) STRANDEDNESS: double	
	(D) TOPOLOGY: circular	
	(ii) MOLECULE TYPE: DNA (genomic)	
25	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
	A LL COTTON COVERED	
20	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
30	(A) ORGANISM: METICODACCCI PYTOTI	
	(ix) FEATURE:	
	(A) NAME/KEY: misc_feature	
	(B) LOCATION 1813	
35	(2) 200200	
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:11:	
	ATGAAAAGT TTGTAGCTTT AGGGCTTCTA TCCGCGGTTT TAAGCTCTTC GTTGTTAGC	C 60
	GAAGGTGATG GTGTTTATAT AGGGACTAAT TATCAGCTTG GACAAGCCCG TTTGAATAG	2 120
40	AATATTTATA ATACAGGGGA TTGCACAGGG AGTGTTGTAG GTTGCCCCCC AGGTCTTAC	C 180
. •	GCTAATAAGC ATAATCCAGG AGGCACCAAT ATCAATTGGC ACTCCAAATA CGCTAATGG	G 240
	GCTTTGAATG GTTTTGGGTT GAATGTGGGT TATAAGAAAT TCTTCCAATT CAAGTCGCT	A 300
•	GATATGACAA GCAAGTGGTT TGGTTTTAGA GTGTATGGGC TTTTTGATTA CGGGCATGC	C 360
	GATTTAGGTA AACAAGTTTA TGCACCTAAT AAAATCCAGT TGGATATGGT CTCTTGGGG	r 420
45	GTGGGGAGCG ATTTGTTAGC TGATATTATT GATAAAGACA ACGCTTCTTT TGGTATTTT	r 480
	GGTGGGGTCG CTATCGGCGG TAACACTTGG AAAAGCTCTG CAGCAAACTA TTGGAAAGA	G 540
	CAAATCATTG AAGCCAAAGG TCCTGATGTT TGTACCCCTA CTTATTGTAA CCCTAATGC	C 600
	CCTTATAGCA CCAACACTTC AACCGTCGCT TTTCAAGTGT GGTTGAATTT TGGGGTGAG	A 660
	GCCAATATCT ACAAGCATAA TGGCGTGGAA TTTGGCGTGA GAGTGCCGCT ACTCATCAA	T 720
50	AAATTTTTGA GCGCGGGTCC TAACGCTACT AACCTTTATT ACCATTTGAA ACGGGATTA	r 780
20	TCGCTTTATT TGGGGTATAA CTACACTTTT TAA	813
	(2) INFORMATION FOR SEQ ID NO:12:	

55 (i) SEQUENCE CHARACTERISTICS:

PCT/US97/19575

	(A) LENGTH: 423 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double	
	(D) TOPOLOGY: circular	
(ii) MOLECULE TYPE: DNA (genomic)	
(iii) HYPOTHETICAL: NO	
(iv) ANTI-SENSE: NO	
	ODICATIVAL COVERCE	
(V1) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
(ix) FEATURE:	
	-(A)-NAME/KEY: misc-feature - (B) LOCATION 1423	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:12:	
ATGCATC	CTA TAATGTTTGC CTATATCGCT AACGCGCTCG CTCAAGCTAG AAAGATCAAC	60
	TTT GCATGGCGTT TCAAAAAATA TCTCAAGTCA AAGAATTAGG CATTGATAAA	120
GCAAAGA	GTT TGATAGGCAA CCTTTCTCAA GTGATTATCT ACCCCACAAA AGATACTGAT	180
	TAG AATGTGGCGT CCCATTAAGC GATAGTGAAA TCAATTTCTT ACACAACACG	240
	GAG CCAGACAAGT GCTAGTAAAA AATATCGTTA CAAACGCTTC AGCTTTTATT	300
	ATT TAAAAAAGAT TTGCAAGAAC TACTTTATAT TCTTGATAGC AATGCTGGTA	360
ATAGAAA TAA	AAT CCTCAATGAT CTTAAAAAAG CAAACCAAGA AACTTATAAG GAAGAGTATT	420 423
(2) INF	ORMATION FOR SEQ ID NO:13:	•
(i)	SEQUENCE CHARACTERISTICS:	
,	(A) LENGTH: 771 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double	
	(D) TOPOLOGY: circular	
_(ii)) MOLECULE TYPE: DNA (genomic)	
(iii) HYPOTHETICAL: NO	
(iv) ANTI-SENSE: NO	
(vi)	ORIGINAL SOURCE:	
•	(A) ORGANISM: Helicobacter pylori	
(ix)) FEATURE:	
	(A) NAME/KEY: misc_feature	
	(B) LOCATION 1771	
(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:13:	
ATGTTGG	GGA GCGTCAAAAA AGCGGTTTTT AGGGTTTTGT GTTTGGGGGC GTTGTGTTTA	60
TGCGGGG	GGT TAATGGCAGA GCAAGATCCT AAAGAGCTTA TATTTTCAGG TATAACTATT	120
	AND ANAROTORS TACACCTARS ANATOTTTC ANAROCTTC CAAATCAAC	180

PCT/US97/19575

- 1 1 ACCCR100 - DML - DOODSIAC

5	GATGCTGATG GCTGTGCAAT CTTAAGAGAG GTTTATTCTA GTGGTAAAGC CATAGCGAGA GAAAACGCAA GAGAGAGCAT TGAAAAAGCT CTTGAACACA CCGCTACTGC TAAAGTTTGT AAATTAAACG ATGCTGAAAA ATGCAAGGAC TTAGCAGAGT TTTATTTTAA TGTAAACGAT CTTAAAAATG CTTTAGAATA TTACTCTAAA TCTTGTAAGT TAAATAATGT TGAAGGGTGT ATGCTGTCAG CAACTTTTTA TAACGATATG ATAAAGGGTT TGAAAAAAGA TAAAAAAGAT CTAGAATATT ATTCTAAAGC TTGCGAGTTA AATAACGGTG GAGGGTGTTC TAAATTAGGA GGGGATTATT TTTTTGGTGA AGGCGTAACA AAAGATTCA AAAAAAGCTTT TGAATATTCT GCCAAAGCTT GTGAGTTGAA CGATGCTAAA GGGTGTTACG CTCTAGCAGC GTTTTATAAT GAGGGTAAAG GCGTGGCAAA GGATGAAAAG CAAACGACAG AAAACCATTA AAAGAGTTGC AAGCTAGGAT TAAAAAGAAC ATGCGATATT CTCAAAGAAC AAAAACCATTA AAAGAGTTGC AAGCTAGGAT TAAAAAGAAGC ATGCGATATT CTCAAAGAAC AAAAACAATA A	240 300 360 420 480 540 600 660 720 771
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 729 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
20	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
25	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Helicobacter pylori	
30	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1729</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:14:	
35	ATGAAAAAT TTTTTTCTCA ATCTTTGTTA GCTCTTATTA TCTCTATGAA TGCGGTATCT GGCATGGATG GTAATGGCGT TTTTTTAGGG GCGGGTTATT TGCAAGGACA GGCGCAAATG CATGCGGATA TTAATTCTCA AAAACAAGCC ACCAACGCTA CGATCAAAGG CTTTGACGCG	60 120 180
40	CTCTTGGGGT ATCAATTTT CTTTGAAAAA CACTTTGGCT TACGCCTTTA TGGGTTTTTT GACTACGCTC ATGCCAATTC TATTAAGCTT AAAAACCCTA ACTATAATAG CGAAGCGGCG CAAGTGGCTA GTCAAATTCT TGGGAAACAA GAAATCAATC GTTTAACAAA CATTGCCGAT CCCAGAACTT TTGAGCCGAA CATGCTCACT TATGGGGGGG CTATGGACGT GATGGTTAAT GTCATCAATA ACGGCATCAT GAGTTTGGGG GCTTTTGGCG GGATACAATT GGCCGGCAAT	240 300 360 420 480 540
45	TCATGGCTTA TGGCGACACC GAGCTTTGAG GGCATTTTAG TGGAACAAGC CCTTGTGAGC AAGAAAGCCA CTTCTTTCCA ATTTTATTC AATGTGGGGG CTCGCTTAAG GATCTTAAAA CATTCTAGCA TTGAAGCGGG CGTGAAATTC CCCATGCTAA AGAAAAACCC CTACATCACT GCAAAAAATT TGGATATAGG GTTTAGGCGC GTGTATTCGT GGTATGTGAA TTACGTGTTC ACTTTCTAG	600 660 720 729
50	(2) INFORMATION FOR SEQ ID NO:15:	
50	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 804 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double	
55	(D) TOPOLOGY: circular	

	(11) MOLECULE TYPE: DNA (genomic)	
5	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
	(vi) ORIGINAL SOURCE:	
10	(A) ORGANISM: Helicobacter pylori	
10	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1804</pre>	
15 .	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:15:	
	A TOTAL COLUMN TOT	٠.
		60 120
		180
20		240
20		300
		360
	•	420
	AGACTTTTAA CGAGCTATCA AATCTTTTTA AACCAAGCCA GAGATAACGC TAACAACCAA	480
25	ATCACAAAAA ACAAAACCCA AAGCCTTGAA GCGATTACAC AAGCTAAAAA CAACGCTAAT	540
	AATGAAATAA GCAACAATCA AACGCAAGCG ATAACTAATA TCACCGAAGC GAAAACGAAC	600
	GCTAATAATG AAATAAGCAA CAATCAAACG CAAGCGATAA CTAACATTAA CGAAGCCAAA	660
	GAAAGCGCTA CAACGCAAAT AAACGCCAAT AAGCAAGAAG CAATAAATAA CATCACGCAA	720
		780
30	ATTGATTTT TTGAGTTTGA ATAA	804
	(2) INFORMATION FOR SEQ ID NO:16:	
	(2) INTOMERITOR TOR DDG ID NO. 20.	
	(i) SEQUENCE CHARACTERISTICS:	
35	(A) LENGTH: 1632 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double	
	(D) TOPOLOGY: circular	
40	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
45	(iv) ANTI-SENSE: NO	
	(vi) ORIGINAL SOURCE:(A) ORGANISM: Helicobacter pylori	
	(ix) FEATURE:	
50	(A) NAME/KEY: misc_feature	
	(B) LOCATION 11632	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:16:	
55	CTCATACACA CCATCCCCAA ACACTCTAAC ATTCTTTTAC CCCCCACACCC CTTTGATACT	60

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	TTAAAAGAGG CGTTTGATAA AATTGACCCC TATACTTTCT TTTTTCCAAA ATTTGAAGCC	120
	ACTAGCACTT CTATTTCTGA TACTAACACG CAGAGGGTGT TTGAAACGCT CAATAACATT	180
	AAAACAAATC TTATAATGAA ATATAGTAAT GAAAATCCAA ACAATTTCAA CACTTGTCCT	240
	TACAATAATA ATGGTAATAC AAAAAATGAT TGTTGGCAAA ATTTCACCCC ACAAACCGCA	300
5	GAAGAATTCA CCAATTTAAT GTTGAACATG ATCGCTGTCT TAGACTCCCA ATCTTGGGGC	360
,	SATGCGATCT TAAACGCTCC TTTTGAATTC ACTAACAGCT CAACAGATTG CGATAGCGAT	420
	CCTTCAAAAT GCGTAAATCC CGGAGTAAAT GGGCGTGTTG ATACTAAAGT CGATCAACAA	480
	PATATACTCA ACAAACAAGG TATTATTAAT AATTTTAGAA AAAAAATAGA AATTGATGCG	540
	STTGTTTTAA AAAATTCAGG GGTTGTAGGG TTAGCCAATG GATATGGCAA TGATGGTGAA	600
10	PATGGCACAT TAGGGGTAGA AGCCTATGCT TTAGATCCTA AAAAACTCTT TGGCAACGAC	660
10	CTTAAGACTA TCAATTTAGA AGATTTAAGA ACCATCTTGC ATGAATTCAG CCACACTAAA	720
	GCTATGGGC ATAACGGGAA TATGACCTAT CAAAGAGTGC CGGTAACGAA AGATGGTCAA	780
	STGGAAAAGG ATAGTAATGG CAAGCCAAAA GATTCTGATG GCCTCCCCTA TAATGTGTGT	840
	CCCTTTATG GGGGATCCAA TCAGCCCGCT TTCCCTAGCA ACTACCCTAA TTCCATCTAT	900
15	CACAATTGTG CGGATGTCCC GGCTGGCTTT TTAGGGGTAA CAGCAGCGGT TTGGCAGCAG	960
1)		020
		080
		140
		.200
20		260
20		320
		380
		440
		500
25		.560
23		.620
		632
30	(2) INFORMATION FOR SEQ ID NO:17: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1071 base pairs	
	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
25	(C) STRANDEDNESS: GOUDIE (D) TOPOLOGY: circular	•
35	(D) TOPOLOGI: CIRCUIAL	
	(ii) MOLECULE TYPE: DNA (genomic)	
40	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Helicobacter pylori	•
45		
	(ix) FEATURE:	
	(A) NAME/KEY: misc_feature	
	(B) LOCATION 11071	*
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:17:	
	TTGATGAAAA GCATTTTGCT CTTTATGATT TTTGTAGTTT GTCAGTTAGA AGGCAAAAAA	60
	TTTTCACAAG ATAATTTTAA GGTGGATTAT AACTACTATT TGCGCAAACA GGATTTGCAC	120
	ATCATTAAAA CGCAAAACGA TTTGTCCAAT GCCTGGTATC TCCCTCCACA AAAAGCCCCC	180
55	AAAGAACATT CTTGGGTGGA TTTTGCTAAA AAATATTTAA ACATGATGGA TTATCTAGGC	240

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	ACTTATTTTT TGCCTTTTTA TCATAGTTTC ACCCCCATTT TTCAATGGTA CCACCCTAAT	300
	ATCAACCCCT ACCAACGCAA TGAGTTTAAG TTCCAAATCA GTTTTAGAGT GCCTGTATTT	360
	AGGCATATTC TTTGGACTAA AGGCACGCTT TATCTGGCTT ATACCCAAAC TAACTGGTTT	420
	CAAATTTATA ATGACCCTCA ATCCGCCCCC ATGCGAATGA TCAATTTCAT GCCTGAACTC	480
5	ATCTATGTTT ATCCTATTAA TTTTAAACCT TTTGGGGGTA AAATAGGGAA TTTTTCTGAA	540
	ATTTGGATAG GTTGGCAGCA CATTTCTAAT GGTGTGGGGG GTGCGCAATG TTACCAGCCT	600
	TTTAATAAAG AAGGTAATCC TGAAAACCAG TTTCCAGGAC AACCTGTAAT CGTTAAAGAT	660
	TATAACGGGC AAAAAGATGT GCGCTGGGGG GGGTGTCKTT CGGTGARCSC GGGCAACSCC	720
	CTGTGTTTCG TTTTGGTGTG GGAAAAGGGA GGCCTAAAAA TCATGGTCGC TTATTGGCCC	780
10	TATGTCCCTT ATGATCAATC CAACCCTCAA TTGATTGATT ACATGGGGTA TGGTAACGCT	840
	AAAATTGATT ACAGGAGAGG GCGCCACCAT TTTGAATTGC AACTTTATGA TATTTTCACG	900
	CAATACTGGC GTTATGATCG CTGGCATGGA GCTTTCCGCT TAGGCTATAC CTACCGCATT	960
	AACCCTTTTG TGGGGATTTA TGCGCAGTGG TTTAACGGCT ATGGCGATGG CTTGTATGAA	1020
	TACGATGTTT TTTCCAATCG TATAGGGGTA GGAATACGCT TGAACCCTTA A	1071
15		
	(2) INFORMATION FOR SEQ ID NO:18:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 2028 base pairs	
20	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double	
	(D) TOPOLOGY: circular	
	(ii) MOLECULE TYPE: DNA (genomic)	
25		
	(iii) HYPOTHETICAL: NO	
	41.)	
	(iv) ANTI-SENSE: NO	
30	(with appearing company	
30	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Helicobacter pylori	
	(ix) FEATURE:	
	(A) NAME/KEY: misc feature	
35	(B) LOCATION 12028	
22	(b) BOCATION 12028	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:18:	
	(AI) DIGODACI DIDOATITION. DIQ ID NO.ID.	
	TTGTCTAAAG GTTTGAGTAT CGGTAATAAA ATCATATTGT GCGTGGCGTT GATTGTGATC	60
40	GTGTGCGTGA GCATTTTAGG GGTGTCCTTA AACAGCAGGG TGAAAGAGAT TTTAAAAGAA	120
	AGCGCTCTGC ATTCAATGCA AGATAGTTTG CATTTCAAGG TTAAGGAAGT GCAAAGTGTT	180
	TTGGAAAACA CTTATACGAG CATGGGCATT GTCAAAGAAA TGCTCCCTGA AGACACCAAA	240
	AGAGAAATCA AAATCCAGTT GTTAAAAAAC TTCATTTTAG CCAATTCGCA TGTCGCTGGG	300
•	GTGAGCATGT TTTTTAAAGA CAGAGAGGAT TTGAGATTGA CGCTTTTACG AGATAACGAT	360
45	ACGATCAAGT TGATGGAAAA CCCGTCATTA GGGAGTAACC CTTTAGCGCA AAAAGCGATG	420
	AAAAATAAAG AAATTTCTAA AAGCTTGCCT TATTACAGGA AAATGCCTAA CGGGGCGGAA	480
	GTTTATGGCG TGGATATTCT TTTACCACTA TTCAAGGAAA ACACGCAAGA AGTGGTGGGG	540
	GTTCTGATGA TTTTCTTTTC CATTGACAGC TTCAGTAATG AAATCACTAA AAACAGGAGC	600
	GATTTATTTT TAATTGGCGT TAAAGGTAAA GTGCTTTTGA GCGCGAATAA AAGCTTGCAA	660
50	GACAAATCCA TCACCGAAAT TTATAAAAGC GTGCCTAAAG CCACTAATGA AGTGATGGCT	720
	ATTTTAGAAA ATGGCTCTAA AGCGACTTTA GAATACTTGG ATCCCTTTAG CCATAAGGAG	780
	AATTTTTTAG CCGTTGAAAC CTTTAAAATG CTAGGCAAAA CAGAAAGTAA AGACAATCTT	840
	AATTGGATGA TCGCTTTGAT CATTGAAAAA GACAAGGTCT ATGAGCAAGT GGGATCGGTG	900
	CGTTTTGTGG TGGTTGCAGC GAGTGCTATC ATGGTGTTAG CCTTAATCAT AGCGATCACT	960
55	CTTTTAATGC GAGCGATCGT GAGCAATCGT TTGGAAGTCG TTTCTAGCAC CTTGTCTCAT	1020

			•		
	TTCTTTAAAT TATTGAACAA TCAAGCO	CAT TCTAGCGACA	TTAAATTGGT	TGAAGCGCGA	1080
	TCTAATGACG AATTAGGGCG CATGCAA	ACA GCGATCAATA	AAAATATCTT	GCAAACCCAA	1140
	AAAACCATGC AAGAAGACAG GCAAGCC	GTC CAAGACACCA	TTAAAGTGGT	TTCAGACGTG	1200
	AAAGCGGGGA ATTTTGCGGT GCGCATC	ACG GCTGAACCCG	CAAGCCCTGA	TTTGAAAGAA	1260
5	TTGAGAGACG CGCTAAATGG GATCATG	GAT TATTTGCAAG	AAAGCGTAGG	GACTCACATG	1320
	CCAAGCATTT TCAAAATCTT TGAAAGC	TAT TCTGGCTTGG	ATTTTAGAGG	GCGGATCCAA	1380
	AACGCTTCGG GTAGGGTGGA ATTGGTT	ACT AACGCTTTAG	GGCAAGAAAT	CCAAAAAATG	1440
	CTAGAAACTT CGTCTAATTT TGCCAAA	GAT CTAGCGAACG	ATAGCGCGAA	TTTAAAAGAA	1500
	TGCGTGCAAA ATTTAGAAAA GGCTTCA	AAC TCCCAACACA	AAAGCCTGAT	GGAAACTTCC	1560
10	AAAACGATAG AAAATATCAC CACTTCC	CATT CAAGGCGTGA	GCTCTCAAAG	TGAAGCCATG	1620
10	ATTGAACAAG GGAAAGACAT TAAAAGC	ATT GTAGAAATCA	TTAGAGATAT	TGCCGATCAA	1680
	ACGAATCTAT TAGCCCTAAA CGCTGCT	TATT GAAGCCGCAC	GAGCCGGCGA	GCATGGCAGA	1740
	GGCTTTGCGG TGGTGGCTGA TGAGGTG				1800
	AGTGAGATTG AAGCCAATAT TAATATT	CTC GTTCAAAGCA	TTTCAGACAC	GAGCGAAAGC	1860
15	ATTAAAAACC AGGTTAAAGA AGTAGAA	GAG ATCAACGCTT	CTATTGAAGC	CTTAAGATCG	1920
	GTTACTGAGG GCAATCTAAA AATCGCT	AGC GATTCTTAG	AAATCAGTCA	AGAAATTGAC	1980
	AAAGTCTCTA ACGATATTTT AGAAGAT				2028
	(2) INFORMATION FOR SEQ ID NO):19:			
20	(2) 1111 01111111111111111111111111111111				
20	(i) SEQUENCE CHARACTERIS	STICS:			
	(A) LENGTH: 816 bas				
	(B) TYPE: nucleic a				
	(C) STRANDEDNESS: C				
25	(D) TOPOLOGY: circu				
23	ν/				
	(ii) MOLECULE TYPE: DNA	(genomic)	•		
(ii) MOLECULE TYPE: DNA (genomic)					
	(II) MODECOLD III . Divi	, 5		•	
	,	, 3 ,			
30	(iii) HYPOTHETICAL: NO	, y =======			
30	,	, 3 - 1 - 2 - 1			
30	(iii) HYPOTHETICAL: NO	, 3 - 1			
30	(iii) HYPOTHETICAL: NO	, 3 - 1 - 2 - 1			
30	(iii) HYPOTHETICAL: NO				
	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE:				
30	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE:				
	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Helic	cobacter pylori			
	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Helic	cobacter pylori _feature			
	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Helic (ix) FEATURE: (A) NAME/KEY: misc	cobacter pylori _feature			
35	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Helic (ix) FEATURE: (A) NAME/KEY: misc	cobacter pylori _feature _6	:		
	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Helic (ix) FEATURE: (A) NAME/KEY: misc (B) LOCATION 181	cobacter pylori _feature 16 N: SEQ ID NO:19			
35	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Helic (ix) FEATURE: (A) NAME/KEY: misc (B) LOCATION 181	cobacter pylori _feature 16 N: SEQ ID NO:19		TTTTAACCTT	60
35	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Helic (ix) FEATURE: (A) NAME/KEY: misc (B) LOCATION 181 (xi) SEQUENCE DESCRIPTION ATGAACATAT TCAAGCGTAT TATTTGG TTAGACGCCA AACACCACAA AGAAAAA	cobacter pylori feature 16 N: SEQ ID NO:19 CGTA ACCGCTATTG	TTTTAGGTTT AAATCACTCG	TGAGCTTAAA	120
35	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Helic (ix) FEATURE: (A) NAME/KEY: misc (B) LOCATION 181 (xi) SEQUENCE DESCRIPTION ATGAACATAT TCAAGCGTAT TATTTGG TTAGACGCCA AACACCACAA AGAAAAA GTGGGCGCTA ACCCTGTGCC GCATGCC	cobacter pylori feature 6 1: SEQ ID NO:19 CGTA ACCGCTATTG AAAA GAAGACCACA GCAA ATCTTGCAAT	TTTTAGGTTT AAATCACTCG CAGTTGTGGA	TGAGCTTAAA TGATTTGAAA	
35	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Helic (ix) FEATURE: (A) NAME/KEY: misc (B) LOCATION 183 (xi) SEQUENCE DESCRIPTION ATGAACATAT TCAAGCGTAT TATTTGG TTAGACGCCA AACACCACAA AGAAAAA GTGGGCGCTA ACCCTGTGCC GCATGCC GAGAAAGGGA TCAAATTAGT GATCGTC	cobacter pylori feature 16 V: SEQ ID NO:19 CGTA ACCGCTATTG AAAA GAAGACCACA GCAA ATCTTGCAAT GTCT TTTACGGATT	TTTTAGGTTT AAATCACTCG CAGTTGTGGA ATGTGTTGCC	TGAGCTTAAA TGATTTGAAA TAATTTAGCG	120
35 40	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Helic (ix) FEATURE: (A) NAME/KEY: misc (B) LOCATION 181 (xi) SEQUENCE DESCRIPTION ATGAACATAT TCAAGCGTAT TATTTGG TTAGACGCA AACACCACAA AGAAAAA GTGGGCGCTA ACCCTGTGCC GCATGCC GAGAAAGGGA TCAAATTAGT GATCGTC CTCAATGACG GCTCTTTAGA CGCGAAA	cobacter pylori feature 16 1: SEQ ID NO:19 CGTA ACCGCTATTG AAAA GAAGACCACA GCAA ATCTTGCAAT GTCT TTTACGGATT	TTTTAGGTTT AAATCACTCG CAGTTGTGGA ATGTGTTGCC GCCCTTATTT	TGAGCTTAAA TGATTTGAAA TAATTTAGCG GGATCGGTTT	120 180 240 300
35 40	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Helic (ix) FEATURE: (A) NAME/KEY: misc (B) LOCATION 181 (xi) SEQUENCE DESCRIPTION ATGAACATAT TCAAGCGTAT TATTTGC TTAGACGCA AACACCACAA AGAAAAA GTGGGCGCTA ACCCTGTGCC GCATGCC GAGAAAGGGA TCAAATTAGT GATCGTC CTCAATGACG GCTCTTTAGA CGCGAAT AATTTGGACA GAAAAATGCA CCTTGTT	feature 16 N: SEQ ID NO:19 CGTA ACCGCTATTG AAAA GAAGACCACA GCAA ATCTTGCAAT GTCT TTTACGGATT TTAC TTCCAGCACC GGGT TTGGCCAATA	TTTTAGGTTT AAATCACTCG CAGTTGTGGA ATGTGTTGCC GCCCTTATTT TCCATGTGGA	TGAGCTTAAA TGATTTGAAA TAATTTAGCG GGATCGGTTT GCCTTTAAGA	120 180 240 300 360
35 40	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Helic (ix) FEATURE: (A) NAME/KEY: misc (B) LOCATION 181 (xi) SEQUENCE DESCRIPTION ATGAACATAT TCAAGCGTAT TATTTGC TTAGACGCA AACACCACAA AGAAAAA GTGGGCGCTA ACCCTGTGCC GCATGCC GAGAAAGGGA TCAAATTAGT GATCGTC CTCAATGACG GCTCTTTAGA CGCGAAT AATTTGGACA GAAAAATCAC AGACAT	feature feature SEQ ID NO:19 GTA ACCGCTATTG AAAA GAAGACCACA GCAA ATCTTGCAAT GTAC TTCCAGCACC GGT TTGCCACTACC GGT TTGCCAATA FAAA AACCTTAAAA	TTTTAGGTTT AAATCACTCG CAGTTGTGGA ATGTGTTGCC GCCCTTATTT TCCATGTGGA AAGGCTCAGT	TGAGCTTAAA TGATTTGAAA TAATTTAGCG GGATCGGTTT GCCTTTAAGA GATTGCTGTG	120 180 240 300 360 420
35 40	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Helic (ix) FEATURE: (A) NAME/KEY: misc (B) LOCATION 181 (xi) SEQUENCE DESCRIPTION ATGAACATAT TCAAGCGTAT TATTTGC TTAGACGCA AACACCACAA AGAAAAA GTGGGCGCTA ACCCTGTGCC GCATGCC GAGAAAGGGA TCAAATTAGT GATCGTC CTCAATGACG GCTCTTTAGA CGCGAAA AATTTGGACA GAAAAATCAC AGACATT TTTTATTCTC AAAAAATCAC AGACATT CCAAATGATC CGGCCAATCA AGGCAGC	cobacter pylori feature 16 N: SEQ ID NO:19 CGTA ACCGCTATTG AAAA GAAGACCACA GCAA ATCTTGCAAT GTCT TTTACGGATT TTAC TTCCAGCACC TGGT TTGGCCAATA TAAA AACCTTAAAA GGCG TTGATTTTAC	TTTTAGGTTT AAATCACTCG CAGTTGTGGA ATGTGTTGCC GCCCTTATTT TCCATGTGGA AAGGCTCAGT TCCATAAACA	TGAGCTTAAA TGATTTGAAA TAATTTAGCG GGATCGGTTT GCCTTTAAGA GATTGCTGTG AGGCCTTATC	120 180 240 300 360
35 40 45	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Helic (ix) FEATURE: (A) NAME/KEY: misc (B) LOCATION 181 (xi) SEQUENCE DESCRIPTION ATGAACATAT TCAAGCGTAT TATTTGC TTAGACGCA AACACCACAA AGAAAAA GTGGGCGCTA ACCCTGTGCC GCATGCC GAGAAAGGGA TCAAATTAGT GATCGTC CTCAATGACG GCTCTTTAGA CGCGAAT AATTTGGACA GAAAAATCAC AGACAT	cobacter pylori feature 16 N: SEQ ID NO:19 CGTA ACCGCTATTG AAAA GAAGACCACA GCAA ATCTTGCAAT GTCT TTTACGGATT TTAC TTCCAGCACC TGGT TTGGCCAATA TAAA AACCTTAAAA GGCG TTGATTTTAC	TTTTAGGTTT AAATCACTCG CAGTTGTGGA ATGTGTTGCC GCCCTTATTT TCCATGTGGA AAGGCTCAGT TCCATAAACA	TGAGCTTAAA TGATTTGAAA TAATTTAGCG GGATCGGTTT GCCTTTAAGA GATTGCTGTG AGGCCTTATC	120 180 240 300 360 420
35 40	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Helic (ix) FEATURE: (A) NAME/KEY: misc (B) LOCATION 181 (xi) SEQUENCE DESCRIPTION ATGAACATAT TCAAGCGTAT TATTTGC TTAGACGCA AACACCACAA AGAAAAA GTGGGCGCTA ACCCTGTGCC GCATGCC GAGAAAGGGA TCAAATTAGT GATCGTC CTCAATGACG GCTCTTTAGA CGCGAAA AATTTGGACA GAAAAATCAC AGACATT TTTTATTCTC AAAAAATCAC AGACATT CCAAATGATC CGGCCAATCA AGGCAGC	cobacter pylori feature 6 N: SEQ ID NO:19 CGTA ACCGCTATTG AAAA GAAGACCACA GCAA ATCTTGCAAT GTCT TTTACGGATT TTAC TTCCAGCACC TGGT TTGGCCAATA TAAA AACCTTAAAA GGCG TTGATTTAC CGCT ACGGAGTTTG	TTTTAGGTTT AAATCACTCG CAGTTGTGGA ATGTGTTGCC GCCCTTATTT TCCATGTGGA AAGGCTCAGT TCCATAAACA ATATTGTCAA	TGAGCTTAAA TGATTTGAAA TAATTTAGCG GGATCGGTTT GCCTTTAAGA GATTGCTGTG AGGCCTTATC AAATCCTTAC	120 180 240 300 360 420 480 540
35 40 45	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Helic (ix) FEATURE: (A) NAME/KEY: misc (B) LOCATION 181 (xi) SEQUENCE DESCRIPTION ATGAACATAT TCAAGCGTAT TATTTGC TTAGACGCCA AACACCACAA AGAAAAA GTGGGCGCTA ACCCTGTGC GCATGCC GAGAAAGGGA TCAAATTAGT GATCGTC CTCAATGACG GCTCTTTAGA CGCGAAA AATTTGGACA GAAAAATGCA CCTTGTT TTTTATTCTC AAAAAATCAC AGACAAT CCAAATGATC CGGCCAATCA AGGCAGC GCTCTCAAAG ACCCCAAGCAA TCTATACC	cobacter pylori feature 6 1: SEQ ID NO:19 CGTA ACCGCTATTG AAAA GAAGACCACA GCAA ATCTTGCAAT TTAC TTTACGGATT TTAC TTCCAGCACC TGGT TTGGCCAATA TAAA AACCTTAAA GGCG TTGATTTTAC CGCT ACGGAGTTTG	TTTTAGGTTT AAATCACTCG CAGTTGTGGA ATGTGTTGCC GCCCTTATTT TCCATGTGGA AAGGCTCAGT TCCATAAACA ATATTGTCAA AGGTTTTAGG	TGAGCTTAAA TGATTTGAAA TAATTTAGCG GGATCGGTTT GCCTTTAAGA GATTGCTGTG AGGCCTTATC AAATCCTTAC GGATGTGGAT	120 180 240 300 360 420 480 540
35 40 45	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Helic (ix) FEATURE: (A) NAME/KEY: misc (B) LOCATION 181 (xi) SEQUENCE DESCRIPTION ATGAACATAT TCAAGCGTAT TATTTGG TTAGACGCCA AACACCACAA AGAAAAA GTGGGCGTA ACCCTGTGC GCATGCC GAGAAAGGGA TCAAATTAGT GATCGTC CTCAATGACG GCTCTTTAGA CGCGAAT TTTTATTCTC AAAAAATCAC AGACATT TTTTATTCTC AAAAAATCAC AGACATT CCAAATGATC CGGCCAATCA AGGCAGC GCTCTCAAAG ACCCAAGCAA TCTATACC AACATCAAAA TCAAACCCCT AGAAGCT	feature 16 1: SEQ ID NO:19 1:	TTTTAGGTTT AAATCACTCG CAGTTGTGGA ATGTGTTGCC GCCCTTATTT TCCATGTGGA AAGGCTCAGT TCCATAAACA ATATTGTCAA AGGTTTTAGG TCACCGGAGC	TGAGCTTAAA TGATTTGAAA TAATTTAGCG GGATCGGTTT GCCTTTAAGA GATTGCTGTG AGGCCTTATC AAATCCTTAC GGATGTGGAT CTTATTTCA	120 180 240 300 360 420 480 540
35 40 45	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Helic (ix) FEATURE: (A) NAME/KEY: misc (B) LOCATION 181 (xi) SEQUENCE DESCRIPTION ATGAACATAT TCAAGCGTAT TATTTGG TTAGACGCCA AACACCACAA AGAAAAA GTGGGCGCTA ACCCTGTGCC GCATGCC GAGAAAGGGA TCAAATTAGT GATCGTC CTCAATGACG GCTCTTTAGA CGCGAAT ATTTATTCTC AAAAAATCAC AGACATT TTTTATTCTC AAAAAATCAC AGACATT CCAAATGATC CGGCCAATCA AGGCAGC GCTCTCAAAG ACCCAAGCAA TCTATACC GGGCTTATCA TAACAGGGAA TCTATACC GGGGCTATCA TCAAACCCCT AGAAGCT GGGGCTATCA TAACAGGGAA TTATGCC	cobacter pylori feature 16 1: SEQ ID NO:19 CGTA ACCGCTATTG AAAA GAAGACCACA GCAA ATCTTGCAAT FITAC TTTACGGATT FITAC TTCCAGCACC FGGT TTGGCCAATA FAAA AACCTTAAAA FGCG TTGATTTTAC FGCT ACGGAGTTTG FGCT TCGCAGCACA CTTG CAAGCAAAAC FCTT GTAGCCTCC	TTTTAGGTTT AAATCACTCG CAGTTGTGGA ATGTGTTGCC GCCCTTATTT TCCATGTGGA AAGGCTCAGT TCCATAAACA ATATTGTCAA AGGTTTTAGG TCACCGGAGC GTGAGGATAA	TGAGCTTAAA TGATTTGAAA TAATTTAGCG GGATCGGTTT GCCTTTAAGA GATTGCTGTG AGGCCTTATC AAATCCTTAC GGATGTGGAT CTTATTTCA TGCGCAAGAT	120 180 240 300 360 420 480 540 600
35 40 45	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Helic (ix) FEATURE: (A) NAME/KEY: misc (B) LOCATION 181 (xi) SEQUENCE DESCRIPTION ATGAACATAT TCAAGCGTAT TATTTGG GAGAAAGGGA TCAAATTAGT GATCGTG CTCAATGACG GCTCTTTAGA CGCGAAT AATTTGGACA GAAAAATCAC AGACATT TTTATTCTC AAAAAATCAC AGACATT CCAAATGATC CGGCCAATCA AGGCAGG GCTCTCAAAG ACCCAAGCAA TCTATAG GCTCTCAAAG ACCCAAGCAA TCTATAG AACATCAAAA TCAAACCCCT AGAAGCT GGGGCTATCA TAACAGGGAA TTATGGC GAAGATAAGG ACTCGCCTTA TGCTAAT	feature 16 1: SEQ ID NO:19 1: SEQ ID	TTTTAGGTTT AAATCACTCG CAGTTGTGGA ATGTGTTGCC GCCCTTATTT TCCATGTGGA AAGGCTCAGT TCCATAAACA ATATTGTCAA AGGTTTTAGG TCACCGGAGC GTGAGGATAA	TGAGCTTAAA TGATTTGAAA TAATTTAGCG GGATCGGTTT GCCTTTAAGA GATTGCTGTG AGGCCTTATC AAATCCTTAC GGATGTGGAT CTTATTTCA TGCGCAAGAT	120 180 240 300 360 420 480 540 600 660 720

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	(2) INFORMATION FOR SEQ ID NO:20:	
. 5	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 486 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
10	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
15	(iv) ANTI-SENSE: NO	
	(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
20	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1486</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:20:	
25	ATGTTTTTA AAACTTATCA AAAATTACTG GGCGCGAGCT GTTTGGCGCT GTATTTAGTG GGCTGTGGGA ATGGTGGTGG CGGTGAATCG CCGGTTGAGA TGATTGCAAA TAGCGAGGGT ACGTTTCAAA TCGACTCCAA AGCAGATAGC ATTACTATTC AAGGCGTGAA GCTTAATAGA GGTAATTGTG CTGTCAATTT TGTTCCAGTA AGTGAGACGT TTCAAATGGG TGTTTTAAGT CAAGTTACTC CAATCTCTAT ACAGGATTTT AAAGATATGG CAAGCACTTA TAAGATATTT	60 120 180 240 300
30	GATCAAAAGA AAGGGTTGGC AAACATAGCA AATAAAATTT CTCAATTAGA GCAAAAGGGT GTGATGATGG AACCTCAAAC CCTTAATTTT GGAGAAAGTT TAAAAGGCAT TTCTCAAGGG TGCAATATTA TAGAGGCAGA AATACAAACC GACAAAGGCG CTTGGACTTT TAACTTTGAT AAATAA	360 420 480 486
35	(2) INFORMATION FOR SEQ ID NO:21:	
40	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1014 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
	(ii) MOLECULE TYPE: DNA (genomic)	
45	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
50	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
55	(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 11014	

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO:21:

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ATGATTAGAT TAAAAGGTTT GAATAAAACT TTAAAAACAA GCTTATTAGC TGGGGTTTTA
                                                                         60
    CTAGGTGCTA CTGCTCCCTT AATGGCAAAG CCTTTATTAA GCGATGAAGA CTTATTGAAA
                                                                        120
    CGAGTAAAAC TACACAATAT CAAAGAAGAT ACGCTGACTA GCTGTAATGC TAAGGTGGAC
                                                                        180
5
    GGCTCTCAAT ACTTGAATAG TGGTTGGAAT TTATCTAAAG AATTTCCGCA AGAATATAGA
    GAAAAGATTT TTGAATGCGT AGAAGAAGAA AAACATAAAC AAGCCCTTAA TTTAATCAAT
    AAAGAAGACA CTGAAGATAA AGAAGAACTT GCAAAAAAAA TCAAAGAAAT TAAAGAAAAA
                                                                        360
    GCTAAAGTTT TAAGGCAAAA ATTTATGGCT TTTGAAATGA AAGAACACTC TAAAGAATTC
                                                                        420
    CCAAATAAAA AGCAACTTCA AACCATGCTT GAGAACGCTT TTGATAATGG AGCTGAAAGT
                                                                        480
10
    TTTATTGATG ATTGGCACGA ACGCTTTGGG GGTATAAGTA GAGAGAATAC TTATAAAGCA
    CTTGGCATTA AAGAATATAG TGATGAAGGA AAGATATTAG CCTTTGGCGA AAGAAGTTAT
    ATTAGACAAT ATAAAAAAGA TTTTGAAGAA AGCACTTATG ATACTAGACA AACCTTATCT
    GCTATGGCTA ATATGAGTGG CGAAAACGAT TATAAAATTA CTTGGTTAAA ACCCAAATAT
    CAGCTCCATA GTTCAAATAA TATTAAACCC TTAATGTCAA ACACAGAGTT GTTAAATATG
    ATAGAGCTAA CCAATATCAA AAAAGAATAT GTTATGGGCT GTAATATGGA AATAGATGGT
                                                                        840
    TCTAAATATC CCATTCATAA AGATTGGGGA TTTTTTGGTA AGGCAAAAGT CCCAGAAACT
                                                                        900
    TGGAGAAATA AGATTTGGGA ATGTATTAAG AATAAAGTAA AGTCCTATGA CAACACTACC
                                                                       960
     GCTGAAATAG GAATAGTTTG GAAAAAAAAT ACTTATTCTA TCTCTCATCA CTAA
                                                                       1014
20
```

(2) INFORMATION FOR SEQ ID NO:22:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1251 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: double
 - (D) TOPOLOGY: circular
- (ii) MOLECULE TYPE: DNA (genomic)
- 30 (iii) HYPOTHETICAL: NO
 - (iv) ANTI-SENSE: NO
- 35 (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Helicobacter pylori
 - (ix) FEATURE:
 - (A) NAME/KEY: misc_feature
- 40 (B) LOCATION 1...1251
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:22:

•	ATGAAAAAAT	TAGTTTTTAG	CATGCTTTTA	TGTTGTAAAA	GCGTGTTTGC	AGAGGGGGAA	- 60
45					AAGCCACTAT		120
					AGGCTCAAGC		180
					CTTTGATTAG		240
					AGTATCAAAT		300
					GGAACGCACA		360
50					TCAATGCTCT		420
50					AAAATGGCGA		480
		ATGTGCAAAA			GCAGTGGCTA		540
		CTGGGGAATT		GCGTGGGGGG	AAATGTTGTG	TAAAATGGTA	600
		ATTATGAAAG		CTTTTAGCAA	CAGGCAATAA	CCCAGAAGAG	660
55				AAAAAGGTTA	ATGATAATAA	GCAGTTAAAA	720

25

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	GATAAACTTG ACCCATTTCT AAAAAGACTT GATGTCCTAC AAACTGAGTT TGGTGTAACT GACCCTACAG CTAACCATAA TAAGCAAGGG ATACATTATT GCACAGAAAA TAAAGAGACA GGTAAATGCG ACCCTATTAA AAATGTATTT AGGACAACTC GCTTAGATAA CGAATTAGAA CAAGAAATCC AAACGCTCAC ACTTGATTTA ATCAAAGCCT CCAATAAAGA CGCTCAAAGC	780 840 900 960
5	CAAGCCTACG CAAATTCAA TCAAAGGATT AAATTACTTA CTCTAAAATA TTTAAAAGAA ATTACCAATC AAATGCTCTT TTTAAATCAA ACAATGGCAA TGCAAAGCGA GATTATGACA GATGATTATT TTAGGCAAAA TAATGATGGC TTTGGGGAAA AAGAAAACCA TATAGACAAA CAATTAACGC AAAAAAGAAT AAACGAAAGA GAAAGAGCTA GAATATACTT TCAAAACCCT AATGTTAAAT TTGACCAATT TGGCTTTCCC ATTTTTAGTA TATGGGATTA A	1020 1080 1140 1200 1251
10	(2) INFORMATION FOR SEQ ID NO:23:	
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1131 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
20	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO	
25	(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
30	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 11131</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:23:	
35	GTGAATAAGT GGATTAAAGG GGCGGTTGTT TTTGTAGGGG GTTTTGCAAC GATTACAACC TTTTCTTTAA TCTACCACCA AAAGCCAAAA GCCCCCCTAA ATAACCAGCC TAGCCTTTTG AATGACGATG AGGTGAAATA CCCCTTACAA GACTACACTT TCACTCAAAA CCCACAGCCA ACTAACACGG AAAGCTCCAA AGACGCTACC ATCAAAGCCT TACAAGAACA GCTCAAAGCC	60 120 180 240
40	GCTTTAAAAG CCCTAAACTC CAAAGAAATG AATTATTCCA AAGAAGAGAC TTTTACTAGC CCTCCCATGG ATCCAAAAAC AACCCCCCCT AAAAAAGACT TTTCTCCAAA ACAATTAGAT TTACTGGCCT CTCGCATCAC CCCTTTCAAG CAAAGCCCTA AAAATTACGA AGAAAACCTG	300 360 420
	ATTTTCCCTG TGGATAACCC TAATGGCATT GATAGTTTCA CTAACCTTAA AGAAAAAGAC ATCGCCACTA ATGAAAACAA GCTTTTACGC ACCATTACAG CTGACAAAAT GATACCCGCT TTTTTGATTA CGCCCATTTC TAGCCAGATC GCTGGTAAAG TGATTGCGCA AGTGGAGAGC	540
45	GATATTTTTG CAAGCATGG CAAAGCCGTC TTAATCCCCA AAGGCTCTAA AGTCATAGGC TATTACAGCA ACAATAACAA AATGGGCGAA TACCGCTTGG ATATTGTATG GAGTCGAATC ATCACTCCCC ATGGCATTAA TATCATGCTC ACTAACGCTA AAGGGGCGGA CATTAAAGGC	660 720
	TATAACGGCT TAGTGGGGGA ATTGATTGAA AGGAATTTCC AACGCTATGG CGTGCCGTTA CTGCTTTCTA CGCTCACTAA CGGCCTATTG ATTGGGATCA CTTCGGCTTT AAACAACAGA	840 900
50	GGCAATAAAG AAGAGGTGAC TAATTTCTTT GGGGATTATC TTTTATTGCA ATTGATGAGG CAAAGCGGCA TGGGGATCAA TCAAGTGGTC AATCAAATTT TAAGAGACAA GAGCAAGATC GCCCCCATTG TGGTGATTAG AGAGGGGAGT AGGGTCTTCA TTTCGCCCAA TACTGACATC	

1131

(2) INFORMATION FOR SEQ ID NO:24:

TTCTTCCCTA TACCCAGAGA GAATGAAGTC ATCGCTGAGT TTTTGAAGTG A

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(i) SEQUENCE CHARACTERISTICS:
              (A) LENGTH: 2751 base pairs
              (B) TYPE: nucleic acid
              (C) STRANDEDNESS: double
               (D) TOPOLOGY: circular
5
        (ii) MOLECULE TYPE: DNA (genomic)
       (iii) HYPOTHETICAL: NO
10
        (iv) ANTI-SENSE: NO
        (vi) ORIGINAL SOURCE:
              (A) ORGANISM: Helicobacter pylori
15
         (ix) FEATURE:
              (A) NAME/KEY: misc feature
               (B) LOCATION 1...2751
         (xi) SEQUENCE DESCRIPTION: SEQ ID NO:24:
20
    GTGGATTTGA GGATCCAATC TAAAGAAGTC AGTCATAATT TAAAGGAATT ATCAAAAACG
                                                                        60
    CTAATCAGCT ATCCTTTTGA AAAACATGTA GAAGCTTTAG GGGAACAATG CAGTAACTTC 120
    GTTTCTATTC CCATTAACAA TGACGACTAT TCAAATATTT GCACTTTTGT GAGTGATTTT 180
    ATAAATCTTA TAGCTTCTTA CAATTTATTA GAATCATTTT TAGATTTTTA TAAAGATAAA 240
    TTAAAATTGA GCGAGCTTGT AACTGAATAT GCCAACGTAA CCAATAATCT GCTTTTCAAA
                                                                       300
    AAATTAATCA AACATTTAAG CGGCAACAAT CAATTGGTTA AAAATTTTTA TCAGTGTATA
                                                                       360
    AGAGAAATTA TAAAATACAA CGCCCCTAAT AAAGAATACA AACCCAATCA ATTTTTTATA
                                                                        420
    ATAGGGAAAG GCAAACAAAA ACAATTAGCA AAAATTTATT CTCATTTAAA AGAACTTAGT
    GCAAGTGAAA TTAAACCACA AGATATGGAA GACATCTTAA AAAAGCTAGA GGAATTAGAT
30
    AAAATTTTTA AAACTACCGA CTTTACAAAA TTCACACCAA AAACTGAAAT TAAGGATATT 600
     ATTAAAGAAA TAGACGAAAA ATACCCTATC AATGAAAATT TTAAACGGCA ATTTAATGAG 660
     TTTGAATCAA ATATTGAAAA ACATGATGAA ATAAAAAAGG ATTTTGAGCG AAACAAAGAG
                                                                       720
     TCGCTGATCC GAGAAATTGA AAATCACTGC AAAAATGAAT GCAATAGCGA AGAAGAGCCG
     GAGTATAAGA TTAATGATCT GCTCAAAAAT ATCCAACAAA TATGCAAAAA TTATATAGAA
35
     AGTCATGCCG TTAATGATGT GTCTAAAGAT ATTAAATCCA TGATGTGTCA GTTTTATTTG
                                                                         900
     AAACAGATAG ATTTATTAGT CAATTCAGAA ATTGTGCGAT ACAGATACAG CAATCTTTTT
     GAACCAATAC AAAGATCTTT ATGGGAGAGT ATAAAAATTT TAGATAATGA AAGTGGCATT
     TATTTGTTCC CTAAAAATAT TGGTGAAATC AAGGATAAAT TTGAAGCAAA CAAGGAAAAA
     TTCAAACAAA GCAAAAATGT TTCTGAGTTC GCAGAATATT GCCGAGAGTG TAACCCCTAT 1140
40
     ACAGCGTTTA ACTTTCATCT AAATATAAAT AATGGTTTAT CTCATCAATT TGAAAAATTC 1200
     GTGCCAATCA TGAAAGAATA CAAAGAGCCA AAAATCACAG ATAATGACCT TGAAGCCATA 1260
     TCAACCAAAG AGACTGGTCT TGCTAGCCAA TTATCTGGGC ACTGGTTTTT TCAGCTTTCG 1320
     TTATTTAATA AAACAAACTT TAATCCTAAT AAAATTTGGA TTCCTTTAGA GTTCAATAAA 1380
     AGATCAAAAA TAAAGTTTGA TAAAGATTTA GAAATCTATT TTGATAGTCA TGAATCGTTC 1440
45
     AATATCTCTA AAAAATACTT GCAAGAAATA GATCAAGAAT CACTAAAAAA GATCAAACAA
     TCAAAAGATT TTTTTCAAT TCAAAAAATA GAGAGTAAGC ATGATAATAA CGATATACTG 1560
     CAACTTGAAT TTTTTGAGAA TGATACAAGT TTTCTTTTTG CTAAAGGAAG TTTTGCAGAA 1620
     ATTTTAGAAT ACAACATGCA ATTAAAAATA GATTCTTTAA TTACAAAAGA ATTTAATAAG 1680
     CTTTTAGCGA TCGTTCAAGA TAGTCCCCAA GATAGTTACC AATTAAAAAT TCGTGTCCGA 1740
50
     CATAACAATA AGCTTCCTAG AGAGAAATAT ACGGAACATG AAATAAAACT TGAAGTTTAT 1800
     GATTGCAGAA AATCCCACGA TCACAATGAG CCAATCATCT TAAGCCAGCA AAGCACCGGC 1860
     TTCCAATGGG CGTTTAATTT CATGTTTGGC TTTCTTTATA ATGTGGGATC ACATTTTAGT
     TTTAACCATA ATATTATCTA TGTCATGGAC GAGCCAGCCA CTCATTTGAG CGTGCCAGCC 1980
   AGAAAGGAGT TTAGGAAATT TTTAAAAGAA TACGCTCATA AAAATCATGT TACTTTTGTT 2040
55
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.5	TTAGCCACCC ATGACCCTT TTTAGTGGAT ACGGATCATT TAGATGAAAT AAGGATTGTG GAAAAGGAAA CAGAAGGCTC TGTAATTAAG AATCACTTTA ACTATCCCCT AAATAATGCA AGCAAAGACT CCGACGCTTT GGACAAAATC AAACGCTCTT TAGGAGTGGG CCAGCATGTT TTTCATAACC CCCAAAAACA CCGAATCATT TTTGTAGAAG GCATCACGGA TTATTGTTAT TTGAGCGCTT TTAAATTGTA TTTGCGTTAC AAAGAATACA AGGACAACCC CATTCCTTTC ACTTTCTTAC CCATTTCAGG GCTTAAAAAC GATTCAAACG ATATGAAAGA AACCATTGAA AAACTTTGCG AGTTAGACAA TCACCCTATT GTTTTGACAG ACGATGACAG AAAATGCGTT TTTAACCAAC AAGCAACGAG CGAACGATT AAAAGAGCTA ATGAAGAAAT GCATGATCCC ATCACCATCC TACAACTCTC AGACTGCGAT AGGCATTTCA AACAAATTGA AGATTGTTTC AGCGCAAACG ATAGAAACAA ATACGCTAAA AATAAGCAAA TGGAATTGAG CATGGCTTTT AAAACAAGGC TTTTGTATGG CGGAGAAGAT GCGATAGAAA AACAAACAAA AAGAAATTTT TTAAAAATTAT TCAAATGGAT TGCATGGGCT ACAAACTTGA TCAAAAACTA A	2100 2160 2220 2280 2340 2400 2520 2520 2580 2640 2700 2751
15	(2) INFORMATION FOR SEQ ID NO:25:	
20	(ii) SEQUENCE-CHARACTERISTICS: (A) LENGTH: 531 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular (ii) MOLECULE TYPE: DNA (genomic)	
25	(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO	
30	<pre>(vi) ORIGINAL SOURCE:</pre>	
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:25:	
40	ATGACTGCAA TGATGCGTTA TTTTCACATC TATGCGACCA CTTTTTTCTT CCCTTTGGCG CTTCTTTTG CGGTTAGTGG GCTTTCATTG CTCTTTAAAG CGCGCCAAGA CACTGGCGCT AAGATCAAAG AATGGGTTT AGAAAAATCC TTAAAAAAAG AAGAACGATT GGACTTTTAA AAAGGCTTTA TAAAAAGAAAA CCATATCGCT ATGCCTAAAA AGATAGAGCC TAGAGAGTAT AGGGGAGCGT TAGTCATTGG CACGCCTTTG TATGAAATCA ACCTTGAAAC TAAAGGCACT CAAACGAAAA TCAAGACCAT TGAAAGGGGC TTTTTAGGCG CGCTCATCAT GCTGCATAAG GCTAAGGTGG GCATCGTGTT TCAGGCGCTT TTAGAGGCAC CTAAACGCAT GTTTATAGGC	60 120 180 240 300 360 420 480
45	GTTTTAATAG GGAGCGTGGT GTTCTTTGGA GCGATCTATT GGTCTTTGTA G	531
50	(2) INFORMATION FOR SEQ ID NO:26: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 669 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular (ii) MOLECULE TYPE: DNA (genomic)	
23	(11) MODECODE TIPE: DNA (GENOMIC)	

	(111) HYPOTHETICAL: NO						
_	(iv) ANTI-SENSE: NO						
5	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>						
10	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1669</pre>						
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:26:						
15	ATGTTTAAAA ACGCTTTAAA TATACAAGAT TTTTCATTTA AAAATCATAC TAGTACAGCC ATTATTGGCA CAAATGGTGC TGGAAAATCA ACGCTTATCA ACACTATTCT AGGCATTAGA	60 120					
	TCAGACTATA ATTTTAAAGC ACAAAACAAT AATATTCCAT ACCACGACAA TGTTATACCA CAACGCAAGC AATTGGGAGT TGTCTCTAAC CTATTCAACT ACCCACCTGG ATTAAACGCA AACGACCTTT TTAAATTCTA TCAATTTTT CACAAAAACT GCACTCTAGA TTTGTTTGAA	180 240 300					
20	AAAAATCTTT TAAATAAAAC CTACGAACAC CTAAGCGACG GACAAAAACA GCGCTTAAAA ATTGACTTAG CTCTTAGCCA TCACCCACAA TTAGTTATTA TGGATGAACC AGAAACCAGT	360 420					
	TTAGAGCAAA ACGCTCTTAT AAGACTATCA AATCTCATAA GCTTGCGCAA CACCCAACAA CTTACAAGTA TCATCGCCAC TCATGATCCT ATTGTCTTAG ATAGTTGCGA ATGGGTATTG CTCCTTAAGA ATGGCAACAT TGCTCAATAC AAACCTTTAA ATTCTATATT AAAATCTGTA	480 540 600					
25	GCTAAAACTT TTAACTTTAA AGAAAAACCA ACCACAAAAG ACTTATTAGC GTTACTAAAG GATATTTAA	660 669					
	(2) INFORMATION FOR SEQ ID NO:27:						
30	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1221 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 						
35	(ii) MOLECULE TYPE: DNA (genomic)						
	(iii) HYPOTHETICAL: NO						
40	(iv) ANTI-SENSE: NO						
40							
	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>						
45	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 11221</pre>						
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:27:						
50	ATGTATGCGG CTCATCCTAT TAAACCCATA AAAGCCCCTA AACTCAAATC TCAATTTTTA AGGCGTGTGT TTGTGGGCGC GTCCATTAGG CGCTGGAATG ACCAAGCATG CCCTTTGGAA TTTGTGGGAAT TAGACAAGCA AGCCCATAAA GCGATGATTG CGTATCTGCT CGCTAAAGAT	60 120 180 240					
55	TTAAAAGATA GGGGTAAAGA TTTAGATTTA GATCTTTTAA TCAAATATTT TTGCTTTGAG TTTTTGGAGC GCTTGGTTTT AACCGATATT AAACCCCCTA TTTTTTACGC CCTCCAACAA	300					

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	ACGCATAGTA AAGAGTTAGC TTCCTATGTT GCGCAAAGTT TGCAAGATGA AATCAGTGCG	360
	TATTTTCTT TAGAGGAACT CAAAGAGTAT TTAAGCCACA GGCCTCAAAT TTTAGAAACT	420
	CAAATTTTAG AGAGCGCGCA TTTTTATGCG TCTAAGTGGG AGTTTGATAT TATCTATCAT	480
	TTTAACCCCA ACATGTATGG CGTGAAAGAG ATTAAAGATA AAATTGACAA GCAACTCCAC	540
5	AATAACGATC ATTTGTTTGA AGGGCTTTTT GGGGAAAAAG AAGATTTGAA AAAATTGGTG	600
_	AGCATGTTTG GGCAGTTGCG TTTCCAAAAG CGCTGGAGCC AAACCCCAAG AGTGCCACAA	660
	ACCAGTGTTC TAGGGCATAC TTTATGCGTG GCGATTATGG GGTATTTATT GAGTTTTGAC	720
	TTGAAAGCTT GTAAAAGCAT GCGGATCAAT CATTTTTTGG GCGGGCTTTT CCATGATTTA	
	CCCGAAATTT TAACCCGAGA CATTATCACG CCCATCAAAC AAAGCGTTGC AGGGCTTGAT	
10	CATTGCATTA AAGAGATTGA AAAAAAGGAA ATGCAAAACA AAGTCTATTC CTTTGTGTCT	
	TTGGGCGTTC AAGAAGATTT GAAATATTTC ACCGAAAACG AGTTTAAAAA CCGCTACAAA	960
	GACAAGTCTC ATCAAATCGT TTTCACTAAA GACGCTGAAG AATTATTCAC GCTTTATAAT	1020
	AGCGATGAAT ATCTTGGGGT TTGCGGGGAG CTTTTGAAGG TGTGCGATCA TTTGAGCGCG	1080
	TTTTTAGAAG CCCAAATCTC TCTTTCTCAT GGCATTTCTA GCTACGATTT AATCCAAGGA	1140
15	GCTAAAAACC TTTTAGAATT GCGATCCCAA ACGGAACTGC TTGATTTGGA TTTAGGGAAA	1200
	-TTGTTTAGAG-ATTTTAAGTA-A	
	(a) TYPODMARTON FOR CEO ID NO.29.	
	(2) INFORMATION FOR SEQ ID NO:28:	
20		
20	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 1008 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double	
	(D) TOPOLOGY: circular	
25		
	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
	(222, 3322	
30	(iv) ANTI-SENSE: NO	
20	(IV) ANII BENEE. NO	
	(vi) ORIGINAL SOURCE:	
	·	
	(A) ORGANISM: Helicobacter pylori	
2.5		
35	(ix) FEATURE:	
	(A) NAME/KEY: misc_feature	
	(B) LOCATION 11008	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:28:	
40		
	GTGTTGTGGG TGCTATATTT TTTAACCAGT TTATTTATTT GCTCTTTGAT TGTTTTGTGG	60
	TCTAAAAAAT CCATGCTCTT TGTGGATAAC GCTAATAAAA TCCAAGGCTT CCATCATGCA	120
	AGAACCCCAC GAGCCGGGGG GCTTGGGATC TTTCTTTCTT TTGCGTTGGC TTGTTATCTT	180
	GAACCTTTTG AGATGCCTTT TAAGGGGCCT TTTGTTTTCT TAGGGCTATC GCTAGTGTTT	240
45	TTGAGCGGTT TTTTAGAAGA CATTAACCTT TCATTAAGCC CCAAAATACG CCTTATTTTG	300
75	CAAGCTGTAG GGGTCGTTTG CATCATTTCA TCAACGCCTT TAGTGGTGAG CGATTTTTCG	360
	CCCCTTTTTA GCTTGCCTTA TTTCATCGCT TTTTTATTCG CTATTTTTAT GCTGGTGGGT	420
	ATCAGTAACG CTATTAATAT CATTGACGGG TTTAACGGGC TTGCATCTGG GATTTGCGCG	480
	ATCGCGCTTT TAGTCATTCA TTATATAGAC CCTAGCAGTT TGTCTTGTTT GCTCGCTTAC	540
50	ATGGTGCTTG GGTTTATGGT GTTAAATTTC CCTTCAGGAA AGATTTTTTT AGGCGATGGG	600
	GGGGCGTATT TTTTGGGTTT GGTGTGCGGG ATTTCTCTCT TGCATTTGAG TTTGGAGCAA	660
	AAAATCAGCG TGTTTTTTGG GCTCAATTTA ATGCTTTATC CGGTCATAGA GGTGCTTTTT	720
	AGTATCCTTA GGCGCAAAAT AAAACGCCAG AAAGCCACCA TGCCGGATAA TTTGCATTTG	780
	CACACCCTTT TATTTAAATT CTTGCAACAA CGCTCTTTCA ATTACCCTAA CCCTTTATGC	840

55 GCGTTTATCC TTATTCTATG CAACCTGCCT TTTATTTTAA TAAGCGTTTT GTTTCGCTTG 900

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	CTTATTTGA ATAGGCAAGT TTGCGCTTTA GAAAAGCGGG CGTTTTAA	.008
5	2) INFORMATION FOR SEQ ID NO:29:	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 291 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double	
10	(ii) MOLECULE TYPE: DNA (genomic)	
15	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
20	(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
20	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1291</pre>	
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:29:	
30	ATGAAAAAGG TTATTGTGGC TTTAGGCGTT TTGGCGTTCG CAAATGTTTT AATGGCAACC SATGTTAAGG CTCTTGTAAA AGGTTGTGCC GCTTGCCATG GGGTTAAGTT TGAAAAGAAA SCTTTAGGTA AAAGCAAAAT CGTTAACATG ATGAGCGAAA AAGAGATTGA AGAGGATCTT ATGGCTTTTA AAAGCGGTGC CAACAAGAAT CCTGTCATGA CCGCGCAAGC TAAAAAAATTA AGCGATGAAG ACATCAAAGC TTTAGCCAAA TACATCCCCA CTCTCAAATA A	60 120 180 240 291
	(2) INFORMATION FOR SEQ ID NO:30:	
35	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 471 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
40	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
45	(iv) ANTI-SENSE: NO	
	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
50	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1471</pre>	
55	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:30:	

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	ATGCGAGATT TCAATAACAT TCAAATCACA CGCTTAAAAG TGCGTCAAAA TGCCGTTTTT	60
	GAAAAACTGG ATCTGGAGTT TAAAGATGGC TTGAGCGCGA TTAGTGGGGC TAGTGGGGTG	120
	GGGAAAAGCG TCCTTATTGC GAGCCTTTTA GGGGCGTTTG GGCTTAAAGA GAGCAACGCT	180
	TCAAACATTG AAGTGGAATT GATCGCGCCT TTTTTAGACA CGGAAGAATA CGGCATTTTT	240
5	AGAGAAGATG AGCATGAACC CTTAGTTATT AGCGTGATTA AAAAAGAAAA AACACGCTAT	300
	TTTTTAAACC AAACAAGCCT ATCTAAAAAC ACGCTCAAAG CGTTATTAAA GGGGCTTATT	360
	AAACGCTTAT CTAACGACAG ATTCAGCCAG AATGAACTCA ACGATATTTT AATGCTCTCC	420
	TTATTAGATG GCTATATCCA AAATAAAAAT AGGCGTTTAG CCCCCTTTTA G	471
10	(2) INFORMATION FOR SEQ ID NO:31:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 357 base pairs	
	(B) TYPE: nucleic acid	
15	(C) STRANDEDNESS: double	
	(D) TOPOLOGY:-circular	
	(ii) MOLECULE TYPE: DNA (genomic)	
20	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
	() OBJECTIVAL COLDER	
25	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
23	(A) ORGANISM: NEITCODACCEI PYTOIT	
	(ix) FEATURE:	
	(A) NAME/KEY: misc feature	
	(B) LOCATION 1357	
30		
	(xi) SEQUENCE DESCRIPTION: SEO ID NO:31:	
	GTGATGCTAA TGGCAATTTT TACCCCTTAT ATTCTTATTT TGAAAATGAT GAAAAAGTCT	60
	ATGAGTTTAT TCGCCAATAT GGGGTTGGAG CAAATTTTTT GCAACAGAGA CATTAAAGAT	120
35	TTAAATGATT TTGTTTTTGG TATAGAAGTG GGGCTTGATA GCAATGCGAG AAAAAATCGT	180
	AGCAGAAAGG CTATGGAAAA TCATCTTATC GGTCTTTTTG TCCAAGCTCA ATTAAATTTT	240
	AAAGAACAAG TAGATATTAG AGAATTTGAG GATTTACGCC AGGCTTTTGG AAATGATACT	300
	AAAAAATTTG ATTTTGTTAT TTTTAGCAAA GAGAAAACTT ATTTTCATAG AAGCTAA	357
40	(2) INFORMATION FOR SEQ ID NO:32:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 1068 base pairs	
45	(B) TYPE: nucleic acid	
43	(C) STRANDEDNESS: double	
	(D) TOPOLOGY: circular	
	(ii) MOLECITE TUDE. DNA (conomic)	
	(ii) MOLECULE TYPE: DNA (genomic)	
50	(iii) HYPOTHETICAL: NO	
50	(III) MIFOIREITCAU. NO	
	(iv) ANTI-SENSE: NO	
	tarry sarah waarware erw	
	(vi) ORIGINAL SOURCE:	
55	(A) ORGANISM: Helicobacter pylori	

(ix) FEATURE:

	(22)	
	(A) NAME/KEY: misc_feature	
	(B) LOCATION 11068	
5		
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:32:	
	ATGAATATCA AAATTTTAAA AATATTAGTT GGAGGGTTAT TTTTTTTGAG CTTGAACGCC	60
	CATTTATGGG GGAAACAAGA CAATAGCTTT TTAGGGATTG GTGAAAGAGC CTATAAAAGC	120
10	GGGAATTATT CTAAAGCGGC GTCTTATTTT AAAAAAGCAT GCAACGATGG GGTGAGTGAA	180
U	GGCTGCACGC AATTAGGAAT CATTTATGAA AACGGGCAAG GCACTAGAAT AGATTATAAA	240
	AAAGCCCTAG AATATTATAA AACCGCATGC CAGGCTGATG ATAGGGAAGG GTGTTTTGGC	300
	TTAGGGGGGC TTTATGATGA GGGTTTAGGC ACGGCTCAAA ATTATCAAGA AGCCATTGAC	360
	GCTTACGCTA AGGCATGCGT TTTAAAACAC CCTGAGAGTT GCTACAATTT AGGCATCATT	420
15	TATGATAGAA AAATCAAAGG CAATGCCGCT CAAGCGGTTA CTTACTATCA AAAAAGCTGT	480
נו	AATTTTGATA TGGCTAAGGG GTGTTATATT TTAGGCACTG CCTATGAAAA AGGCTTTTTA	540
	GAAGTCAAAC AGAGCAACCA TAAAGCCGTT ATCTATTATT TGAAAGCGTG CCGATTGAAT	600
	GAGGGGCAGG CTTGCCGAGC GTTAGGGAGT TTGTTTGAAA ATGGCGATGC AGGGCTTGAT	660
	GAGGGGCAGG CIIGCCGAGC GIIACGGAGI IIGIIIGIGII IIGGGCTTTAAA CAATTCTGGT	720
20	GAAGATITIG AAGIGGCGIT TOATTATTO CITTETOTOTO OF TOATTAAAAA AGACCCCCAA	780
20	AAGGCTTTTA ACTATTTCAA GCAAGCATGC GATATGGGGA GCGCGGTGAG TTGCTCTAGG	840
	ATGGGCTTTA TGTATTCGCA AGGGGACACT GTTTCAAAAG ACTTGAGGAA AGCCCTTGAT	900
	AATTATGAAA GAGGTTGCGA TATGGGCGAT GAAGTGGGTT GCTTCGCTCT AGCGGGCATG	960
	TATTACAACA TGAAAGATAA AGAAAACGCC ATAATGATTT ATGACAAGGG CTGTAAATTG	1020
25	GGCATGAAAC AGGCATGCGA AAATCTCACC AAACTCAGGG GGTATTAG	1068
23	OGCATORATE MODELLE VALUE CANADA CANAD	
	(2) INFORMATION FOR SEQ ID NO:33:	
	(i) SEQUENCE CHARACTERISTICS:	
30	(A) LENGTH: 582 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double	
	(D) TOPOLOGY: circular	
2.5	(ii) MOLECULE TYPE: DNA (genomic)	
35	(11) MOLECULE TYPE: DNA (Genomic)	
	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
40		
	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Helicobacter pylori	
	(ix) FEATURE:	
45	(A) NAME/KEY: misc_feature	
	(B) LOCATION 1582	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:33:	
50	ATGAAAGAAA AAAACTTTTG GCCTTTAGGA ATCATGAGCG TGCTTATTTT TGGGCTTGGG	60
50	ATGAGAGAAA AAAACTITIG GCCTTTAGGA ATCATGAGCG TGCTTATTT TGGGGTTGTAT	120
	TTCAAGGGTC ATAACGAAGT GGATTTAAAC TTTAACGCCA TGCTTAAAAC TTATGAAAAC	180
	TTTAAATCCA ATTATCGTTT TTCAGTGGGT TTAAAGCCTC TTACCGAAAG CCCTAAAACC	
	CCCATTTTGC CCTATTTTC TAAAGGCACG CATGGGGATA AAAAAATCCA AGAAAACCTT	300
55	TTAAACAACG CTTTGATTTT AGAAAAGTCC AACACGCTTT ATGCACAATT GCAACCGCTC	
رر	TIMAMCANCE CITIENTITI NORMANOTO MICROCATTI ATGCACANTI GGILLOGOT	

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(iii) HYPOTHETICAL: NO

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AAACCCGCTT TAGATTCGCC AAATATTCAA GTGTATTTAG CGTTCTATCC CAGCCAATCC CAGCCCAGAT TATTAGGAAC GCTTGATTGT AAAAACGCAT GCGAACCTTT AAAATTTGAT TTGTTAGAGG GCGATAAAGT GGGGCGCTAT AAGATCCTTT TTAAATTTGT TTTTAAAAAT AAAGAAGAAT TGATTTTGGA GCAACTGGCT TTTTTTAAGT AG	420 480 540 582
(2) INFORMATION FOR SEQ ID NO:34:	
 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 870 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
(ii) MOLECULE TYPE: DNA (genomic)	
(iii) HYPOTHETICAL: NO	
(iv) ANTI-SENSE: NO	
(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1870</pre>	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:34:	
AGCGCCTTAC GCTTTTATGG GGAATATTTA GGGGGGGCGA TGAAAGGGTT TAAAAGCGAT TCTTTAGCTT CTTATCAAAC CGCAAGCTTG AATATTGATC TGTTGATGGA TAAGCCTATT GACAAAGAAA AAAGGTTTGC GTTAGGGATA TTTGGAGGCG TTGGAGTGGG GTGGAATGGG ATGTATCAAA ATTTAAAAGA GATTAGAGGG TATTCACAGC CTAACGCCTT TGGGTTGGTG TTAAATTTAG GGGTGAGCAT GACGCTCAAC CTCAAACACC GCTTTGAATT AGCCCTAAAA ATGCCTCCCT TAAAAGAAAC TTCGCAAACC TTTTTATATT ATTTTAAAAG CACTAATATT TATTATATTA GTTACAACTA TTTATTGTAA	60 120 180 240 300 360 420 480 540 660 720 780 840 870
(2) INFORMATION FOR SEQ ID NO:35: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 2007 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular (ii) MOLECULE TYPE: DNA (genomic)	

(iv) ANTI-SENSE: NO

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Helicobacter pylori

(ix) FEATURE:

(A) NAME/KEY: misc_feature

(B) LOCATION 1...2007

10

5

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:35:

	ATGAGAAAAC	TATTCATCCC	ACTTTTATTA	TTCAGCGCTT	TAGAAGCGAA	CGAGAAAAAC	60
	CCCTTTTTCA	TAGAAGCCGG	CTTTGAAACT	GGGCTATTAG	AAGGCACACA	AACGCAAGAA	120
15	AAAAGACACA	CCACCACAAA	AAACACTTAC	GCAACTTACA	ATTATTTACC	CACAGACACG	180
13	ATTTTAAAAA	GAGCGGCTAA	TTTATTCACC	AATGCCGAAG	CGATTTCAAA	ATTAAAATTC	240
	TCATCTTTAT	CCCCTGTTAG	AGTGTTGTAT	ATGTATAATG	GTCAATTAAC	TATAGAAAAC	300
	THE PROPERTY OF THE PROPERTY O	ATAATTTAAA	TAATGTTAAG	CTTAGTTTTA	CAGACGCTCA	AGGCAACACG	360
	ATTGATCTAG	GCGTGATAGA	GACCATCCCC	AAACACTCTA	AGATTGTTTT	ACCCGGGGAG	420
20	CCCTTTGATA	GTTTAAAAGA	GGCGTTTGAT	AAAATTGACC	CCTATACTTT	ATTTCTTCCA	480
20	AAATTTGAAG	CCACTAGCAC	TTCTATTTCT	GATACTAACA	CGCAGAGGGT	GTTTGAAACG	540
	CTCAATAACA	TTAAAACAAA	TCTTATAATG	AAATATAGTA	ATGAAAATCC	AAACAATTTC	600
	AACACTTGTC	CTTACAATAA	TAATGGTAAT	ACAAAAAATG	ATTGTTGGCA	AAATTTCACC	660
	CCACAAACCG	CAGAAGAATT	CACCAATTTA	ATGTTGAACA	TGATCGCTGT	CTTAGACTCC	720
25	CAATCTTGGG	GCGATGCGAT	CTTAAACGCT	CCTTTTGAAT	TCACTAACAG	CTCAACAGAT	780
	TGCGATAGCG	ATCCTTCAAA	ATGCGTAAAT	CCCGGAGTAA	ATGGGCGTGT	TGATACTAAA	840
	GTCGATCAAC	AATATATACT	CAACAAACAA	GGTATTATTA	ATAATTTTAG	AAAAAAATA	900
	GAAATTGATG	CGGTTGTTTT	AAAAAATTCA	GGGGTTGTAG	GGTTAGCCAA	TGGATATGGC	960
	AATGATGGTG	AATATGGCAC	ATTAGGGGTA	GAAGCCTATG	CTTTAGATCC	TAAAAAACTC	1020
30	TTTGGCAACG	ACCTTAAGAC	TATCAATTTA	GAAGATTTAA	GAACCATCTT	GCATGAATTC	1080
50	AGCCACACTA	AAGGCTATGG	GCATAACGGG	AATATGACCT	ATCAAAGAGT	GCCGGTAACG	1140
	AAAGATGGTC	AAGTGGAAAA	GGATAGTAAT	GGCAAGCCAA	AAGATTCTGA	TGGCCTCCCC	1200
	татаатстст	GTTCGCTTTA	TGGGGGATCC	AATCAGCCCG	CTTTCCCTAG	CAACTACCCT	1260
	AATTCCATCT	ATCACAATTG	TGCGGATGTC	CCGGCTGGCT	TTTTAGGGGT	AACAGCAGCG	1320
35	GTTTGGCAGC	AGCTCATCAA	TCAAAACGCC	TTGCCGATCA	ACTACGCTAA	CTTGGGGAGT	1380
	CAAACAAACT	ACAACCTAAA	CGCTAGTTTA	AACACGCAAG	ATTTAGCCAA	TTCCATGCTC	1440
	AGCACCATCC	AAAAAACCTT	TGTAACTTCT	AGCGTTACCA	ACCACCATTT	TTCAAACGCA	1500
	TCGCAAAGTT	TTAGAAGCCC	TATTTTAGGG	GTTAACGCTA	AAATAGGCTA	TCAAAACTAC	1560
	TTTAATGATT	TCATAGGGTT	GGCTTATTAT	GGCATCATCA	AATACAATTA	CGCTAAAGCT	1620
40	GTTAATCAAA	AAGTCCAGCA	ATTGAGCTAT	GGTGGGGGGA	TAGATTTGTT	ATTGGATTTC	1680
40	ATCACCACTT	ACTCCAATAA	AAATAGCCCT	ACAGGCATTC	AAACCAAAAG	GAATTTTTCT	1740
		GTATCTTTGG			ACAGCTATTA	TGTGTTGAAC	1800
		GAAGCGGCAA		GCTACCGGGT		CTATAAGCAT	1860
•	TCTAAATATT	CTGTAGGGAT	TAGCATCCCT	TTAATCCAAA		CGTCGTTTCT	1920
45	AGCGGTGGCG	ATTATACGAA	CTCTTTTGTT	TTCAATGAAG	GGGCTAGCCA	CTTTAAGGTG	1980
		ACGGGTGGGT				•	2007

- (2) INFORMATION FOR SEQ ID NO:36:
- 50 (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 192 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: double
 - (D) TOPOLOGY: circular

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	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
5	(iv) ANTI-SENSE: NO	
	(vi) ORIGINAL SOURCE:(A) ORGANISM: Helicobacter pylori	
10	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1192</pre>	•
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:36:	
20	ATGAATACAG AAATTTTAAC CATCATGTTA GTTGTCTCCG TGCTTATGGG ATTGGTAGGC- TTAATAGCGT TTTTATGGGG GGTTAAAAGC GGTCAGTTTG ACGATGAAAA ACGCATGCTT GAAAGCGTGT TGTATGACAG CGCGAGCGAC TTGAACGAAG CGATTTTACA AGAAAAACGC CAAAAGAATT AA	-60 120 180 192
20	(2) INFORMATION FOR SEQ ID NO:37:	
25	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1221 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
30	(ii) MOLECULE TYPE: DNA (genomic) (iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
35	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
40	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 11221</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:37:	
45	ATGGTATTTT TTCATAAGAA AATTATTTTA AATTTTATCT ATTCTTTAAT GGTTGCTTTT TTATTCCATT TATCCTATGG GGTTCTTTTA AAAGCCGATG GAATGGCTAA AAAGCAAACT CTTTTAGTGG GTGAAAGGCT TGTGTGGGAT AAGCTCACGC TGTTAGGGTT TTTAGAAAAA AACCATATCC CCCAAAAACT CTACTACAAT TTGAGCTCTC AAGATAAAGA ATTGAGTGCT	60 120 180 240
50	GAAATCCAAA GCAATGTTAC CTACTACACT TTAAGAGATG CAAATAACAC GCTCATTCAA GCCCTTATCC CTATTAGCCA GGATTTGCAA ATCCATATTT ACAAAAAAGG AGAGGATTAT TTTTTAGACT TTATCCCCAT TGTTTTCACT CGTAAAGAAA GAACCCTCCT TCTTTCCTTA CAAACTTCGC CCTATCAAGA TATTGTCAAA GCCACCAATG ACCCCCTTTT AGCCAACCAA TTGATGAACG CGTATAAAAA AAGCGTGCCT TTTAAACGCC TAGTGAAAAAA CGATAAAATC	300 360 420 480 540
55	GCTATCGTTT ATACAAGGGA TTATCGTGTG GGGCAAGCGT TTGGCCAGCC GACCATCAAA ATGGCGATGG TTAGCTCTCG TTTGCACCAA TACTATCTTT TTTCCCATTC AAACGGCGT TATTACGATT CAAAAGCGCA AGAAGTGGCA GGGTTTTTAC TAGAAACCCC GGTGAAATAC	600 660 720

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	ACCCGCATTT CTTCGCCTTT TTCGTATGGG AGGTTCCATC CTGTTTTAAA AGTTAAACGG	780					
	CCTCATTACG GCGTGGATTA TGCGGCTAAA CATGGCAGTT TGATCCATTC TGCTTCAGAC	840					
	GGCCGTGTGG GTTTTATAGG GGTTAAGGCG GGTTATGGGA AGGTGGTTGA AATCCATTTG	900					
	AATGAATTGC GCTTGGTGTA TGCTCACATG AGCGCGTTCG CTAACGGATT AAAAAAAAGGC	960					
_	TCGTTCGTTA AAAAAGGGCA AATCATAGGA AGAGTGGGAA GCACGGGTTT AAGCACCGGG	1020					
5	CCGCATTTGC ATTTTGGCGT GTATAAAAAC TCCCGCCCCA TTAATCCTTT AGGCTATATC	1080					
	CCGCATTTGC ATTTTGCCGT GTATAAAAAC TCCCGCCCCA TTAATCCTTT AGGCTATTCAGC CGCACCGCTA AAAGCAAGCT GCATGGCAAA CAAAGAGAGG TTTTTTTAGA AAAAGCTCAG	1140					
	CGCACCGCTA AAAGCAAGCT GCATGGCAAA CAAAGAGAGG TITTTTTAGA AAAACTCAG TATTCTAAGC AAAAATTAGA AGAACTTTTT AAAACCCATT CTTTTGAAAA AAATTCATTT	1200					
		1221					
	TATCTTTTAG AGGGTTTTTA A	1221					
10	TO TO NO 20						
	(2) INFORMATION FOR SEQ ID NO:38:						
	CONTRACT CUADACTED ICHICC.						
	(i) SEQUENCE CHARACTERISTICS:						
	(A) LENGTH: 891 base pairs						
15	(B) TYPE: nucleic acid						
	(C) STRANDEDNESS: double						
	(D) TOPOLOGY: circular						
	(ii) MOLECULE TYPE: DNA (genomic)						
20	· · · · · · · · · · · · · · · · · · ·						
	(iii) HYPOTHETICAL: NO						
	(iv) ANTI-SENSE: NO						
	(IV) ANTI-SENSE: NO						
25	(vi) ORIGINAL SOURCE:						
25	(A) ORGANISM: Helicobacter pylori						
	(A) ORGANISM. METICODACTEL PYTOLI						
	(ix) FEATURE:						
	(A) NAME/KEY: misc_feature						
30	(B) LOCATION 1891						
50	(b) Bootilett British						
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:38:						
	TTGTTTTTAG TCAAAAAAT AGGCGTGGTA ATAATGATTT TAGTCTGCTT TTTAGCTTGC	60					
35	TCGCAAGAGA GCTTTATCAA AATGCAAAAA AAAGCCCAAG AGCAAGAAAA TGACGGCTCT	120					
<i></i>	AAACGCCCCA GCTATGTGGA TTCGGATTAT GAAGTCTTTA GCGAAACGAT TTTTTTACAA	180					
	AACATGGTGT ATCAGCCTAT AGAGGAAAGA AACGCTTTTT TCCAACTGAC TAAAGATGAA	240					
	GACAATTCTT TTAACCCTGA AAATTCCGTG ATTTTACTGA ATGAGCCAAG CGATAATAGT	300					
	GAAAAAACC TACTCTCATA CCCAAACGAT CCCAATAACA ATGAAGACAA CGCTAATAAT	360					
40	AGTCAAAAAA ATCCGTTCCT TTACAAGCCC AAAAGAAAAA CAAAAAAACCC AAAACTCATT	420					
	GAATATTCCC AACAAGATTT CTACCCCCTA AAAAATGGGG ATATTATCAT GAGTAAAGAA	480					
	GGGGATCAAT GGTTGATAGA AATCCAATCC AAAGCCTTGA AGCGTTTTTT AAAAGATCAA	540					
	AACGATAAAG ATCGCCAGAT CCAAACTTTC ACTTTTAATG ACACTAAAAC GCAAATCGCG	600					
·	CAAATTAAGG GCAAAATTTC TTCGTATGTT TATACCACCA ATAACGGTAG CTTGAGTTTA	660					
45	AGGCCTTTTT ATGAATCGTT TTTGTTAGAA AAAAAGAGCG ATAATGTTTA TACGATAGAG	720					
	AATAAGGCTT TAGATACTAT GGAGATTTCA AAGTGTCAAA TGGTGTTAAA AAAGCATTCA	780					
	ACCGATAAAT TAGACAGCCA GCATAAAGCC ATCAGTATTG ATTTGGATTT TAAAAAAAGAG	840					
	CGCTTTAAGA GCGATACGGA ACTCTTTTTA GAATGTCTTA AGGAAAGTTA G	891					
	, 						
50	(2) INFORMATION FOR SEQ ID NO:39:						
23	· · · · · · · · · · · · · · · · · · ·						
	(i) SEQUENCE CHARACTERISTICS:						
	(A) LENGTH: 747 base pairs						
	(B) TYPE: nucleic acid						
55	(C) STRANDEDNESS: double						

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	(D) TOPOLOGY: circular	
	(ii) MOLECULE TYPE: DNA (genomic)	
5	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
	(vi) ORIGINAL SOURCE:	
10	(A) ORGANISM: Helicobacter pylori	
	(ix) FEATURE:	
	(A) NAME/KEY: misc_feature	
	(B) LOCATION 1747	
15	(xi)_SEQUENCE_DESCRIPTION:_SEQ_ID_NO:39:	
	GTGAGCTATG ACAACACCGA TGATTATTAT TTCCCTAGAA ATGGGGTTAT CTTTAGTTCC	60
	TATGCGACAA TGTCTGGTTT GCCAAGCTCT GGCACGCTCA ATTCTTGGAA CGGGTTAGGC	120
20	GGGAATGTCC GTAACACCAA AGTTTATGGT AAATTCGCCG CTTACCACCA TTTGCAAAAA	180
	TATTTATTGA TAGATTTGAT CGCTCGTTTT AAAACGCAAG GGGGCTATAT CTTTAGGTAT	240 300
	AACACCGATG ATTACTTGCC CTTAAACTCC ACTTTCTACA TGGGGGGCGT AACCACGGTG AGAGGCTTTA GGAACGGCTC AATCACACCT AAAGATGAGT TTGGCTTGTG GCTTGGAGGC	360
	GATGGGATTT TTACCGCTTC TACTGAATTG AGCTATGGGG TGTTAAAAGC GGCTAAAATG	420
25	CGTTTAGCGT GGTTTTTTGA CTTTGGTTTC TTAACCTTTA AAACCCCAAC TAGGGGGAGT	480
	TTCTTCTATA ACGCTCCCAC CACGACGGCG AATTTTAAAG ATTATGGCGT TGTAGGGGCT	540
	GGGTTTGAAA GGGCGACTTG GAGGGCTTCT ACAGGCTTAC AGATTGAATG GATTTCGCCC	600
	ATGGGGCCTT TGGTGTTGAT TTTCCCTATA GCGTTTTTCA ACCAATGGGG CGATGGCAAT	660
	GGCAAAAAAT GTAAAGGGCT GTGCTTTAAC CCTAACATGA ACGATTACAC GCAACATTTT	720
30	GAATTTTCTA TGGGAACAAG GTTTTAA	747
	(2) INFORMATION FOR SEQ ID NO:40:	
	(i) SEQUENCE CHARACTERISTICS:	
35	(A) LENGTH: 1008 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double	
	(D) TOPOLOGY: circular	
40	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
45	(iv) ANTI-SENSE: NO	
	(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
	(ix) FEATURE:	
50	(A) NAME/KEY: misc_feature	
	(B) LOCATION 11008	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:40:	
55	GTGCAACACT TCAATTTCCT CTATAAAGAT TCTTTATTTT CTATCGCTTT ATTCACTTTC	60

900

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ATTATCGCTC TTGTGATTTT ATTAGAACAG GCTAGAGCGT ATTTCACCCG AAAGAGAAAC
    AAAAAATTTT TGCAAAAATT CGCCCAAAAT CAAAACGCCT ATGCGAGCAG CGAGAATTTA
                                                                        180
    GACGAGCTTT TAAAGCATGC TAAAATTTCC AGTTTGATGT TTTTAGCTAG GGCGTATTCT
                                                                        240
    AAAGCGGATG TGGAAATGAG CATTGAAATC TTAAAAGGGC TTTTGAATCG CCCCTTAAAA
                                                                        300
    GATGAAGAAA AAATCGCTGT TTTAGATTTA TTGGCTAAAA ATTATTTTAG CGTGGGGTAT
                                                                        360
5
    TTGCAGAAAA CAAAAGACAC CGTGAAAGAA ATTTTGCGCT TTTCCCCAAG GAATGTGGAA
                                                                        420
    GCGTTGTTGA AGCTTTTGCA TGCGTATGAA TTAGAAAAAG ATTATTCAAA GGCTTTAGAA
                                                                        480
    ACTTTGGAAT GTTTGGAAGA ATTAGAGGTG CCTAAAATTG AAACGATTAA AAATTACCTC
                                                                        540
    TATTTAATGC ATTTAATAGA GAATAAGGAA GATGCGGCTA AAATCTTGCA TGTTTCAAAA
                                                                        600
    GCGTCGTTAG ATTTGAAAAA AATCGCTCTG AATCACTTAA AATCGCATGA TGAAAATCTT
                                                                        660
10
    TTTTGGCAAG AAATTGATAC AACCGAACGG CTAGAAAATG TGATCGATCT TTTATGGGAT
                                                                        720
    ATGAATATCC CTGCTTTTAT TTTAGAAAAA CATGCCCTTT TGCAGGACAT CGCGCGATCT
                                                                        780
    CAAGGGTTGC TTTTGGATCA CAAACCTTGC CAAATTTTTG AATTAGAGGT TTTACGCGCT 840
    CTATTGCATA GCCCTATAAA AGCGAGTCTG ACTTTTGAAT ACCGCTGCAA GCATTGCAAA
                                                                        900
    CAAATCTTTC CTTTTGAAAG CCATAGGTGT CCTGTGTGTT ACCAGTTAGC GTTTATGGAT 960
                                                                       1008
    ATGGTGCTTA AAATCTCTAA AAAAACGCAT GCTATGGGAG TGGATTAA
     (2) INFORMATION FOR SEQ ID NO:41:
         (i) SEQUENCE CHARACTERISTICS:
20
               (A) LENGTH: 1242 base pairs
               (B) TYPE: nucleic acid
               (C) STRANDEDNESS: double
               (D) TOPOLOGY: circular
25
        (ii) MOLECULE TYPE: DNA (genomic)
        (iii) HYPOTHETICAL: NO
       (iv) ANTI-SENSE: NO
30
         (vi) ORIGINAL SOURCE:
             (A) ORGANISM: Helicobacter pylori
         (ix) FEATURE:
35
              (A) NAME/KEY: misc feature
              (B) LOCATION 1...1242
         (xi) SEQUENCE DESCRIPTION: SEQ ID NO:41:
40
     ATGAGGAAAA TTTTTTCTTA TATTTCTAAG GTTCTATTAT TTATTGGGGT GGTTTATGCA
     GAGCCTGATT CTAAAGTGGA AGCCTTAGAA GGGAGGAAGC AAGAGTCTTC TTTGGATAAA 120
     AAAATCCGCC AAGAATTGAA GAGTAAGGAA TTGAAGAATA AGGAATTAAA GAATAAGGAT
                                                                       180
     TTGAAAAATA AAGAAGAAAA GAAAGAAACA AAAGCCAAGA GAAAACCCAG AGCAGAAGTC
     CATCATGGGG ACGCCAAAAA TCCCACTCCA AAGATCACGC CTCCTAAAAT CAAAGGGAGT
45
     AGTAAGGGCG TTCAAAATCA AGGCGTTCAA AACAACGCGC CAAAACCTGA AGAAAAAGAT 360
    ACAACCCCTC AAGCTACTGA AAAAAATAAG GAAACAAGCC CTAGCTCTCA ATTCAATTCC 420
     ATTTTTGGTA ATCCTAATAA CGCTACCAAC AACACCCTTG AAGATAAGGT CGTAGGGGGC
    ATTTCATTGC TTGTTAATGG TTCGCCTATC ACGCTGTATC AAATCCAAGA AGAGCAAGAA
    AAATCTAAAG TGAGTAAGGC TCAAGCTAGG GATCGTTTGA TCGCTGAACG CATTAAAAAC
                                                                        600
50
     CAAGAAATTG AGCGCTTAAA AATCCATGTA GATGATGACA AGCTAGACCA AGAAATGGCG
                                                                         660
     ATGATGGCGC AACAACAAGG CATGGATTTA GACCATTTCA AACAGATGCT TATGGCTGAG
     GGGCATTATA AACTCTATAG AGATCAACTT AAAGAGCATT TAGAAATGCA AGAATTGTTG
                                                                         780
    CGTAATATTT TGCTCACGAA TGTGGATACC AGCTCTGAAA CCAAAATGCG CGAATATTAC
                                                                         840
```

AACAAACACA AGGAGCAATT CAGTATCCCC ACAGAAATAG AAACCGTGCG CTACACTTCC

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	ACCAATCAAG AGGATTTAGA AAGGGCTATG GCAGACCCTA ATTTGGAAGT CCCAGGGGTG AGTAAGGCCA ATGAAAAAAT AGAGATGAAA ACCCTAAACC CTCAAATCGC CCAAGTCTTT ATTTCGCATG AGCAAGGCTC TTTCACGCCC GTTATGAATG GGGGTGGGGG GCAGTTCATC ACCTTTTATA TCAAGGAAAA AAGGGGTAAA AATGAAGTGA GCTTCAGTCA GGCCAAGCAA	960 1020 1080 1140
. 5	TTCATCGCCC AAAAATTAGT GGAAGAATCT AAGGATAAGA TTTTAGAAGA GCATTTTGAA AAATTGCGCG TTAAGTCTAG GATTGTGATG ATCAGAGAGT GA	1200 1242
	(2) INFORMATION FOR SEQ ID NO:42:	
10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 561 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: circular	
15	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
20	(iv) ANTI-SENSE: NO	
	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
25	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1561</pre>	
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:42:	
	ATGATTAAAA GAATTGCTTG TATTTTAAGC TTGAGCGCGA GTTTAGCGTT AGCTGGCGAA GTGAATGGGT TTTTCATGGG TGCGGGTTAT CAACAAGGTC GTTATGGCCC TTATAACAGC AATTACTCTG ATTGGCGTCA TGGCAATGAC CTTTATGGTT TGAATTTCAA ATTAGGTTTT	60 120 180
35	GTAGGCTTTG CCAATAAATG GTTTGGGGCT AGGGTGTATG GCTTTTTAGA TTGGTTTAAC ACTTCAGGGA CTGAACACAC CAAAACCAAT TTGCTCACCT ATGGCGGCGG TGGCGATTTG ATTGTCAATC TCATTCCTTT GGATAAATTC GCTCTAGGTC TCATTGGTGG CGTTCAATTA	300 360
	GCCGGAACA CTTGGATGTT CCCTTATGAT GTCAATCAAA CCAGATTCCA GTTCTTATGG AATTTAGGCG GAAGAATGCG TGTTGGGGAT CGCAGTGCGT TTGAAGCGGG CGTGAAATTC	420 480
40	CCTATGGTTA ATCAGGGTAG CAAAGATGTA GGGCTTATCC GCTACTATTC TTGGTATGTG GATTATGTCT TCACTTTCTA G	540 561
	(2) INFORMATION FOR SEQ ID NO:43:	
	(i) SEQUENCE CHARACTERISTICS:	
45	(A) LENGTH: 729 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: circular	
50	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
6.5	(iv) ANTI-SENSE: NO	
55	· · · · · · · · · · · · · · · · · · ·	

ICDOCID: JAIO 081832381 I

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	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
5	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1729</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:43:	
10	ATGAAAAAT TTTTTCTCA ATCTTTGTTA GCTCTTATTA TCTCTATGAA TGCGGTATCT	60
	CCCATGGATG GTAATGGCGT TTTTTTAGGG GCGGGTTATT TGCAAGGACA GGCGCAAATG	120
	CATGCGGATA TTAATTCTCA AAAACAAGCC ACCAACGCTA CGATCAAAGG CTTTGACGCG	180
	CTCTTGGGGT ATCAATTTTT CTTTGAAAAA CACTTTGGCT TACGCCTTTA TGGGTTTTTT	240 300
	GACTACGCTC ATGCCAATTC TATTAAGCTT AAAAACCCTA ACTATAATAG CGAAGCGGCG	360
15	CAAGTGGCTA GTCAAATTCT TGGGAAACAA GAAATCAATC GTTTAACAAA CATTGCCGAT CCCAGAACTT TTGAGCCGAA CATGCTCACT TATGGGGGGG CTATGGACGT GATGGTTAAT	420
	CCCAGAACTT TTGAGCCGAA CATGCTCACT TATGGGGGGG CTATGGACGT GATGGTTATA GTCATCAATA ACGGCATCAT GAGTTTGGGG GCTTTTGGCG GGATACAATT GGCCGGCAAT	480
	TCATGGCTTA TGGCGACACC GAGCTTTGAG GGCATTTTAG TGGAACAAGC CCTTGTGAGC	540
	AAGAAAGCCA CTTCTTTCCA ATTTTTATTC AATGTGGGGG CTCGCTTAAG GATCTTAAAA	600
20	CATTCTAGCA TTGAAGCGGG CGTGAAATTC CCCATGCTAA AGAAAAACCC CTACATCACT	660
20	GCAAAAAATT TGGATATAGG GTTTAGGCGC GTGTATTCGT GGTATGTGAA TTACGTGTTC	720
	ACTTCTAG	729
	(2) INFORMATION FOR SEQ ID NO:44:	
25		
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 771 base pairs (B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double	
30	(D) TOPOLOGY: circular	
50	(2)	
	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
35		
	(iv) ANTI-SENSE: NO	
	A LA COMPANY COMPANY	
	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
40	(A) ORGANISM: NEITCODACCCI PYTOIT	
40	(ix) FEATURE:	
	(A) NAME/KEY: misc_feature	
	(B) LOCATION 1771	
	TO TO NO. 44	
45	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:44:	
	ATGGGATACG CAAGCAAATT AGCTTTAAAG ATTTGTTTGG TAGGTTTATG TTTATTTAGC	60
	ACCCTTGGTG CAGAACACCT TGAGCAAAAA GGGAATTATA TTTATAAGGG AGAGGAGGCT	120
	TATAATAATA AGGAATATGA GCGAGCGGCT TCTTTTTATA AGAGCGCTAT TAAAAATGGT	180
50	GAGTCGCTTG CTTATATTCT TTTAGGGATC ATGTATGAAA ATGGTAGGGG TGTACCTAAA	240
	GATTACAAGA AAGCGGTTGA ATATTTCCAA AAAGCTGTTG ATAACGATAT ACCTAGAGGG	300
	TATAACAATT TGGGCGTGAT GTATAAAGAG GGTAAGGGAG TTCCTAAAGA TGAAAAGAAA	360
	GCGGTGGAAT ATTTTAGAAT AGCTACAGAG AAAGGTTATA CTAACGCTTA TATCAACTTA	420
	GGCATCATGT ATATGGAGGG CAGGGGAGTT CCAAGTAACT ATGCGAAAGC GACAGAATGT	480
55	TTTAGAAAAG CGATGCATAA GGGCAATGTG GAAGCTTATA TTCTCCTAGG GGATATTTAT	540

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TATAGCGGGA ATGATCAATT GGGTATTGAG CCGGACAAAG ATAAGGCTGT TGTCTATTAT AAAATGGCGG CTGATGTGAG TTCTTCTAGA GCTTATGAAG GGTTGTCAGA GTCTTATCGG 660 TATGGGTTAG GCGTGGAAAA AGATAAAAAA AAGGCTGAAG AATACATGCA AAAAGCATGC 720 GATTTTGACA TTGATAAAAA TTGTAAGAAA AAGAACACTT CAAGCCGATA A 771 5 (2) INFORMATION FOR SEQ ID NO:45: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1974 base pairs (B) TYPE: nucleic acid 10 (C) STRANDEDNESS: double (D) TOPOLOGY: circular (ii) MOLECULE TYPE: DNA (genomic) 15 (iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO 20 (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori (ix) FEATURE: (A) NAME/KEY: misc feature 25 (B) LOCATION 1...1974 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:45: ATGAGAAAAC TATTCATCCC ACTTTTATTA TTCAGCGCTT TAGAAGCGAA CGAGAAAAAC GGCTTTTCA TAGAAGCCGG CTTTGAAACT GGGCTATTAG AAGGCACACA AACGCAAGAA 120 30 AAAAGACACA CCACCACAAA AAACACTTAC GCAACTTACA ATTATTTACC CACAGACACG ATTTTAAAAA GAGCGGCTAA TTTATTCACC AATGCCGAAG CGATTTCAAA ATTAAAATTC TCATCTTTAT CCCCTGTTAG AGTGTTGTAT ATGTATAATG GTCAATTAAC TATAGAAAAC TTCTTGCCTT ATAATTTAAA TAATGTTAAG CTTAGTTTTA CAGACGCTCA AGGCAATGTG 420 35 ATCGATCTAG GCGTGATAGA GACTATCCCC AAACACTCTA AGATTGTTTT GCCCGGAGAG 480 GCATTTGATA GTCTAAAAAT TGACCCCTAT ACTTTATTTC TTCCAAAAAT TGAAGCCACT AGCACTTCTA TTTCTGACGC TAACACGCAG AGGGTGTTTG AAACGCTCAA TAAGATTAAG 540 ACAAATTTGG TCGTAAATTA TAGGAATGAA AACAAATTTA AAGATCACGA AAATCATTGG 600 GAAGCCTTTA CCCCACAAAC CGCAGAAGAA TTCACTAATT TAATGTTGAA CATGATCGCT 660 GTTTTAGACT CCCAATCTTG GGGCGATGCG ATCTTAAACG CTCCTTTTGA GTTCACTAAC 720 40 AGCCCAACAG ATTGCGATAA TGATCCTTCA AAATGCGTAA ATCCTGGGAC AAACGGGCTT GTCAATTCTA AAGTCGATCA AAAATATGTG TTAAACAAAC AAGACATTGT CAATAAATTT 840 900 AAAAACAAAG CGGATCTTGA TGTAATTGTT TTAAAGGATT CAGGGGTTGT AGGGCTTGGG AGTGATATTA CCCCTAGCAA CAATGATGAT GGCAAGCATT ATGGCCAGTT AGGGGTAGTA GCTTCTGCTT TAGATCCTAA AAAACTCTTT GGCGATAACC TTAAGACTAT CAATTTAGAG 1020 45 GATTTAAGAA CCATCTTGCA TGAATTCAGC CACACTAAAG GCTATGGGCA TAACGGGAAT ATGACCTATC AAAGAGTGCC GGTAACGAAA GATGGTCAAG TGGAAAAGGA TAGTAATGGC 1200 AAGCCAAAAG ATTCTGATGG CCTCCCCTAT AATGTGTGTT CGCTTTATGG GGGATCCAAT CAGCCCGCTT TCCCTAGCAA CTACCCTAAT TCCATCTATC ACAATTGTGC GGATGTCCCG 1260 50 GCTGGCTTTT TAGGGGTAAC AGCAGCGGTT TGGCAGCAGC TCATCAATCA AAACGCCTTG 1320 CCGATCAACT ACGCTAACTT GGGGAGTCAA ACAAACTACA ACCTAAACGC TAGTTTAAAC 1380 ACGCAAGATT TAGCCAATTC CATGCTCAGC ACCATCCAAA AAACCTTTGT AACTTCTAGC 1440 GTTACCAACC ACCATTTTC AAACGCATCG CAAAGTTTTA GAAGCCCTAT TTTAGGGGTT 1500 AACGCTAAAA TAGGCTATCA AAACTACTTT AATGATTTCA TAGGGTTGGC TTATTATGGC 1560 55 ATCATCAAAT ACAATTACGC TAAAGCTGTT AATCAAAAAG TCCAGCAATT GAGCTATGGT 1620

5	GGGGGGATAG ATTIGITATI GGATTICATC ACCACTATION OF ACC	1680 1740 1800 1860 1920 1974
10	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 504 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
15	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
20	(iv) ANTI-SENSE: NO	
	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
25	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1504</pre>	
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:46:	
30	ATGAAATTGG TGAGTCTTAT TGTAGCGTTA GTTTTTTGTT GTTTTTTAGG GGCTGTAGAG TTGCCTGGAG TTTATCAAAC TCAAGAATTT TTATACATGA AAAGCTCTTT TGTGGAGTTT TTTGAGCATA ACGGGAAGTT CTATGCCTAT GGTATTTCTG ATGTGGATGG CTCTAAAGCC AAAAAAAGACA AACTCAATCC TAACCCAAAG CTAAGGAATC GCAGCGATAA AGGCGTGGTG	60 120 180 240
35	TTTTTAAGCG ATTTGATTAA GGTTGGGGAA CAATCTTATA AAGGCGGTAA GGCGTATAAT TTTTATGACG GCAAGACCTA CCATGTGAGA GTCACTCAAA ATTCAAACGG GGATTTGGAA TTCACTTCAA GCTATGACAA ATGGGGGTAT GTGGGCAAAA CCTTCACCTG GAAACGCCTG AGCGATGAAG AAATCAAAAA TCTAAAGCTC AAGCGTTTTA ACTTGGACGA AGTCCTTAAA ACCCTCAAAG ATAGCCCTAT TTAA	300 360 420 480 504
40		
	(2) INFORMATION FOR SEQ ID NO:47:	
	(i) SEQUENCE CHARACTERISTICS:	
45	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 885 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
	(A) LENGTH: 885 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double	
45 50	(A) LENGTH: 885 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: circular	
	(A) LENGTH: 885 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular (ii) MOLECULE TYPE: DNA (genomic)	

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(A) ORGANISM: Helicobacter pylori (ix) FEATURE: (A) NAME/KEY: misc feature 5 (B) LOCATION 1...885 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:47: ATGAGTAATC AAGCGAGCCA TTTGGATAAT TTTATGAACG CTAAAAATCC CAAAAGTTTT 60 TTTGATAATA AGGGGAATAC CAAATTCATC GCTATCACAA GCGGTAAGGG GGGCGTGGGG 10 AAATCCAACA TTAGCGCTAA TTTAGCTTAC TCTTTATACA AGAAAGGTTA TAAGGTAGGG 180 GTATTTGATG CGGATATTGG TTTAGCGAAT TTAGATGTCA TTTTTGGGGT GAAAACCCAT 240 AAAAATATCT TGCATGCCTT AAAAGGCGAA GCCAAATTGC AAGAAATCAT TTGCGAGATT GAACCCGGGC TTTGCTTAAT CCCTGGGGAT AGCGGCGAAG AAATTTTAAA ATACATCAGC GGCGCGGAAG CTTTGGATCG ATTCGTAGAT GAAGAGGGGG TTTTAAGCTC TTTAGATTAT 15 420 ATTGTGATTG_ATACGGGTGC_TGGGATTGGG_GCCACTACGC_AAGCGTTTTT_GAATGCGAGC---480---GATTGCGTGG TGATTGTTAC CACACCCGAT CCTTCAGCGA TTACCGATGC GTATGCATGC ATTAAAATCA ACTCCAAGAA TAAAGATGAA TTGTTCCTTA TCGCTAACAT GGTAGCCCAA 600 CCTAAAGAAG GĈAGGGCĜAC TTATGAAAGG CTATTCAAGG TGGCTAAAAA CAATATCGCT 660 TCATTAGAAT TGCACTATTT AGGGGCGATT GAAAACAGCT CCTTATTGAA ACGCTATGTG 20 720 AGGGAGCGAA AGATTTTGAG GAAAATAGCC CCTAACGATT TGTTTTCGCA ATCCATTGAC CAGATAGCGA GCCTTTTAGT TTCTAAACTA GAAACCGGCA CTTTAGAAAT ACCAAAAGAA 840 GGTTTAAAAA GCTTTTTTAA AAGGCTTTTG AAGTATTTGG GGTAG 885 25 (2) INFORMATION FOR SEQ ID NO:48: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1119 base pairs (B) TYPE: nucleic acid 30 (C) STRANDEDNESS: double (D) TOPOLOGY: circular (ii) MOLECULE TYPE: DNA (genomic) 35 (iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: 40 (A) ORGANISM: Helicobacter pylori (ix) FEATURE: (A) NAME/KEY: misc feature (B) LOCATION 1...1119 45 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:48: TTGGAACCTT CAAGAAATCG CCTAAAACAT GCCGCCTTTT TTGTGGGGCT TTTTATCGTT TTGTTTTAA TTATAATGAA GCACCAAACC TCCCCCTATG CTTTCACGCA TAATCAAGCC 120 50 CTTGTCACTC AAACCCCCCC CTATTTCACG CAACTCACTA TCCCTAAACC AAATGACGCT 180 TTAAGCGCGC ATGCGAGCTC TTTAATCAGC TTGCCTAACG ACAATCTTTT GAGCGCTTAT 240 TTTAGCGGCA CTAAAGAAGG GGCAAGGGAT GTGAAAATCA GCGCGAATCT TTTTGACAGC 300 AAGACTAATC GCTGGAGCGA AGCCTTCATT CTTTTAACCA AAGAAGAGCT TTCTCATCAT 360

TCGCATGAAT ACATCAAAAA ATTAGGTAAC CCCTTGCTTT TTTTGCATGA TAATAAAATT

TTGTTGTTTG TCGTAGGGGT GAGCATGGGC GGGTGGGCCA CTTCTAAAAT CTATCAATTT

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	GAAAGCGCTT TAGAGCCGAT TCATTTTAAG TTTGCGCGAA AACTCTCTTT AAGCCCTTTT	540
	TTAAATTTGA GCCATTTAGT AAGGAATAAG CCTTTAAACA CCACTGATGG CGGGTTTATG	600
	CTACCACTCT ATCACGAATT AGCCACCCAA TACCCCTTGT TGTTGAAATT TGACCAACAA	660
	AATAACCCAA GAGAGCTTTT AAGGCCTAAT ACCTTAAACC ACCAGCTCCA ACCAAGCTTA	720
5	ACCCCCTTTA AAGACTGCGC TGTCATGGCG TTTAGAAACC ATTCTTTTAA AGATAGCCTC	780
-	ATGCTAGAAA CCTGTAAAAC CCCCACTGAT TGGCAAAAAC CCATTTCTAC AAATCTTAAA	840
	AACTTAGATG ATTCTTTAAA TTTACTCAAT TTAAATGGAA TATTGTATTT GATCCACAAC	900
	CCTAGCGATT TATCACTGCG TCGTAAAGAA CTTTGGCTTT CTAAATTAGA AAACTCCAAC	960
	TCGTTTAAAA CCTTAAAAGT TTTGGATAAA GCGAATGAAG TGAGTTACCC AAGCTATAGC	1020
10	CTTAATCCGC ATTTTATAGA TATTGTCTAT ACTTACAACC GCTCTCATAT CAAACACATC	1080 1119
	CGTTTCAATA TGGCTTATTT AAATTCCCTT CTCAAGTGA	1119
	AND COMPANY TON TON ONE TO NO. 40.	
	(2) INFORMATION FOR SEQ ID NO:49:	
15	(i) SEQUENCE CHARACTERISTICS:	
13	(A) LENGTH: 2937 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double	
	(D) TOPOLOGY: circular	
20		
	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
25	() THE GRACE NO	
25	(iv) ANTI-SENSE: NO	
	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Helicobacter pylori	
30	(ix) FEATURE:	
	(A) NAME/KEY: misc_feature	
	(B) LOCATION 12937	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:49:	
35		60
	ATGAAGAAA GAAAACATGT ATCCAAGAAA GTGTTTAATG TCATTATCTT GTTTGTGGCA GTATTCACTC TTTTAGTCGT CATTCACAAA ACCCTTTCAA ACGGCATTCA CATACAAAAT	120
	TTAAAAATTG GAAAACTTGG CATTCTGAA TTATACTTAA AACTCAATAA CAAGCTTTCT	180
	TTGGAAGTTG AGCGGGTTGA TCTCTCTTCT TTCTTCCATC AAAAACCCAC TAAAAAGCGT	240
40	TTAGAAGTTT CTGATTTGAT TAAAAATATC CGTTATGGCA TTTGGGCGGT GTCTTATTTT	300
40	GAAAAACTTA AAGTCAAAGA AATCATTTTA GACGATAAAA ATAAAGCCAA TATCTTTTTT	360
	GATGGGAATA AATACGAGTT AGAATTTCCA GGAATCAAAG GGGAATTTTC CCTAGAAGAC	420
	GATAAAAATA TCAAGCTTAA AATCATCAAT TTGCTTTTTA AAGATGTTAA AGTCCAAGTG	480
	GATGGCAACG CCCACTATTC ACCCAAAGCC AGGAAAATGG CGTTCAATTT GATTGTCAAG	540
45	CCCTTAGTTG AACCCAGCGC TGCAATTTAT TTGCAAGGGC TAACCGATTT AAAAACCATA	600
	GAATTAAAAA TTAACACTTC TCCAATGAAA AGCCTAGCGT TTTTAAAGCC TCTTTTCCAA	660
	CGCCAATCGC AAAAAAATTT AAAAACGTGG ATTTTTGACA AGATCCAATT TGCCAGCTTT	720
	AAGATTGATA ACGCTTTAAT CAAGGCTAAT TTCACTCCTA GCGAGTTTAT CCCATCGCTT	780
	TTGGAAAATT CTGTAGTTAA AGCCACTTTG ATTAAGCCTT CAGTCGTTTT TAATGATGGC	840 900
50	TTATCGCCCA TTAAAATGGA TAAAACCGAA TTGATTTTCA AAAACAAACA GCTCCTCATA	960 960
	CAGCCCCAAA AAATCACTTA TGAAACCATG GAATTAACCG GCTCTTACGC CACTTTTTCC	1020
	AATTTGTTAG AAGCCCCTAA GTTGGAGGTT TTTTTAAAAA CGACCCCTAA TTATTATGGC GATAGCATTA AGGATTTATT GAGCGCTTAT AAAGTCGTTT TACCTTTGGA TAAAATCAGC	1020
	ATGCCATCTA GCGCGGATTT GAAGCTCACT TTGCAATTCT TAAAAAAACAC CGCCCCCTTA	1140
55	ATGCCATCTA GCGCGGATTT GAAGCTCACT TIGCAATICT TAAAAAACAC CGCCCCCT	

TTTAGCGTTC AAGGCAGCGT TAATTTGCAA GAAGGCACTT TCTCGCTCTA TAATATCCCC 1200

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	•	
	CTTTACACGC AAAGCGCTCA AATCAATTTG GACATCGCCC AAGAATACCA ATACATCTAC	1260
	ATAGACACGA TCCACACGCG CTATGCAAAC ATGCTGGATT TAGACGCTAA AATCGCTTTA	1320
	GATTTAGGTC AAAAAAACCT TTCTTTGGAT TCTTTAGTCC ATAAAATCCA AGTCAATACC	1380
	AATAACAATA TCAACATGCG CTCTTATGAT CCCAATAACA CTCAAGAAGA TCCGCAAACT	1440
5	AACTTTACTT TGGATCTAAA AAGCTTGCAT TCTATCATTC AAGAGGGTGA AAATTCAGAA	1500
	GTTTTTAGAA GAAAAATCAT AGACACCATT AAAGCCCAAA GCGAAGATAA ATTCACTAAA	1560
	GATGTTTTTT ACGCCACAGG AGACACTCTC AAAAGCCTGT CGTTGAGTTT TGATTTTTCC	1620
	AACCCCGATC ACATACAATG GAGCGTGCCA CAACTCTTAT TAGAAGGCGA ATTTAAAGAT	1680
	AACGCCTATA CTTTTAAGAT CAAAGATTTG AAAAAGATCA AGCCCTATTC CCCCATTATG	1740
10	GACTATATTG CCCTAAAAGA CGGCTCTTTA GAGGTTTCTA CGAGCGATTT TGTCAATATT	1800
10	GATTTTTTG CTAAAGATTT GAAAATCAAC CTCCCCATTT ATAGGAGCGA TGGATCGCAT	1860
	TTTGATTCTT TTTCTTTATT TGGCTCTATC AATAAAGATG AAATTTCTGT CTATACTCCA	1920
	AGCAAAAGCA TATCCATAAA AGTTAAGGGG GATCAAAAGG ATATTACCCT TAATAACATT	1980
	GATTTGAGTA TTGATGATTT CTTGGATAGT AAAATGCCAG CTATTGCGGG ATTATTCTCA	2040
15	AAAGAACGAA AAGAAAAGCC TAGCTCTAAA GAAATCCAAG ATGAAGATGT TTTCATTAGC	2100
13	•	2160
	-GCCAAACAAC-GCTATGAAAA-AGCCCAGAAA-ATTATCCCCA-TCTCTACACG-CATCCATGCT	
	AAAGATGTCG TGCTGATCTA TAAAAAAATG CCTTTTCCTT TAGAAAATCT TGATATTGTC	2220
	GCTCAAGACG ATAGGGTGAA AATTGATGGC AATTATAAAA ACGCCATGAT CATGGCGGAT	2280
30	TTAGTGCATG GGGCTTTGTA TCTTAAGGCT CATAATTTTA GCGGGGATTA TATCAACACC	2340
20	ATTCTTCAAA AAGATTTCGT AGAAGGAGGC TTATTCACGC TTATTGGGGC TCTTGAAGAT	2400
	CAGGTTTTCA ATGGCGAATT GAAATTCCAA AACACAAGCT TAAAGAATTT CGCCCTCATG	2460
	CAAAACATGG TCAATCTCAT CAACACCATT CCCTCCCTCA TTGTCTTTAG AAACCCTCAT	2520
	TTAGGGGCTA ATGGCTATCA AATCAAAACC GGCTCCGTTG TGTTTGGGAT CACTAAAGAA	2580
25	TATTTAGGGT TAGAAAAAAT TGATCTTGTC GGCAAAACGC TTGATATTGC TGGCAATGGA	2640
25	ATCATTGAAT TAGACAAAAA CAAATTAGAT TTAAACTTAG AAGTTTCCAC TATCAAGGCT	2700
	TTGAGTAATG TCTTAAATAA AATCCCTATC GTGGGCTATC TCGTTTTAGG AAAAGGAGGT	2760
	AAAATCACCA CTAACGTGAA TGTCAAAGGC ACGTTGGATA AGCCTAAAAC CCAAGTAACT	2820
	TTAGCGTCAG ATATTATCCA AGCGCCTTTT AAAATCTTAC GCCGTATTTT CACGCCTATT	
20	GACATCATCG TGGATGAAGT CAAGAAAAAC ATTGATTCAA AAAGGAAATT AAAATGA	2937
30		
	(2) INFORMATION FOR SEQ ID NO:50:	
	(i) SEQUENCE CHARACTERISTICS:	
25	(A) LENGTH: 1434 base pairs	
35	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double	
	(D) TOPOLOGY: circular	
4.0	(ii) MOLECULE TYPE: DNA (genomic)	
40		
	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
45	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Helicobacter pylori	
	(ix) FEATURE:	
	(A) NAME/KEY: misc_feature	
50	(B) LOCATION 11434	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:50:	
	ATGAATACTA TTATAAGATA TGCGAGTTTA TGGGGCTTGT GTATTACTCT AACTCTAGCG	60
55	CAAACCCCCT CTAAAACCCC TGATGAAATC AAGCAAATCC TTAACAATTA TAGCCATAAG	120

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	AATTTAAAGC TCATTGATCC	GCCGACAAGT	TCTTTAGAAG	CGACACCGGG	TTTTTTACCC	180
	TCGCCTAAAG AAACAGCGAC					240
	GATAAAGCCG CTTTGGGGCT	TTATGAATTG	CTAAAGGGGG	CTACCACCAA	TCTCAGTTTG	300
	CAAGCGCAAG AACTCAGTGT					360
5	TTTTTGCCTA CTTTGAACGC	GAGTTATAAT	TTTAAAAATG	AAGCTAGGGA	TACTCCAGAA	420
	TATAAGCATT ATAACACCCA					480
	TTTAGCAATG TGAATAATGT					540
	TTAGAATATA GCCGCCAAAG					600
	AACAATCTCG CTCGCATGAT					660
10	AAAAGGGTTA CTAAGCTCTA					720
•	AAAGCGCAAG GGAATTTGAG					780
	AACCGCTTGA CTTTAGAATA					840
	ATTGATGCGC CTAATTTGCA					900
	ATTTCTGCAC TCAGATACCA					960
15	GACTCATGGC TTTTTTGGAT					1020
13	TACCCAGGTC AGCAAAATAC					1080
	GGGTTGAGCT TGCAAAAACA					1140
	GCGTATAAAA AATTGGAGCA					1200
	GCCAGAGCTA AGATTGAATC					1260
20	AATATTAAAA GGAAATACGA					1320
20	ACCACGCGCT TTGATGCAGA					
	AAAGCCAATT ACATTTTTAA					1434
	(2) INFORMATION FOR SI	EQ ID NO:51	:			
25	(=,	-				
	(i) SEQUENCE CHAI	RACTERISTICS	S:			
	(A) LENGTH:				•	
	(B) TYPE: n					
	(C) STRANDE	ONESS: doub	le			
30	(D) TOPOLOG					
•						
	(ii) MOLECULE TYP	E: DNA (geno	omic)			
	,,	•				
	(iii) HYPOTHETICAL	: NO				
35	• •					
	(iv) ANTI-SENSE: I	70				
	(= - /					
	(vi) ORIGINAL SOU	RCE:				
	(A) ORGANISI	M: Helicoba	cter pylori			
40	• •					
, ,	(ix) FEATURE:					
	(A) NAME/KE	Y: misc feat	ture			
	(B) LOCATION					
45	(xi) SEQUENCE DESC	CRIPTION: SI	EQ ID NO:51	:		
	·					
	ATGCTATCTT TTATAAGCGC	GTTTGATAAA	AGGGGCGTTT	CAATACGCCT	TCTAACAGCC	60
	TTGTTACTGC TTTTTAGTTT					120
	AAATACCTTT CTAAAAATCA					180
50	TCTCAAGAAA AAGTCGTTAG					240
50	GCTAACGTGA GCGATTTTTT					300
	TTGTCTCAAA AAGTGGATTT					360
	GAAAACAAA AAAAAATATT					420
	ATGATAAACG GCATTGAAAA					480
55	ATTAAAAATT TAGAAAACAC					540
			-			

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	GCGATCGCCA AGTTAGAAAT TTTAAAATCG CTATTAGAAA TCCAAAAAAA CGATTTAGAA GTAGCGCTCT CTAGCAGCCA TTATTCCATG GGCGAATTGA CTTTTAAAGA AAACGAGATT	600 660
	TTAAGCATTG CCCCTAAAAA TTTTGAATTC AATAACGAGC AAGAGCTGCA TAACATTAGC	720
		780
_	GCCACTAATT ACGATATTGC GATCGCCAGG CTTGATGAAG AAAAAGCACA AAAAGACATC	
5	ACTCTGGCTA AAAAAGCTT TTTAGAAGAC ATAAACGTTA CCGGGGTGTA TTATTTCCGC	840
	TCCAAACAAT ACTATAACTA CGACATGTTT AGCGTCGCTT TGTCTATCCC TTTACCTCTT	900
	TATGGCAAGC AGGCTAAATT AGTGGAGCAA AAGAAAAAAG AAAGCTTGGC GTTTAAAAGC	960
	GAAGTGGAAA ACGCCAAAAA CAAAACGCGC CACCTGGCCC TAAAACTCCT TAAAAAATTA	1020
	GAAACCTTGC AAAAAAACCT GGAATCGATC AATAAAATCA TCAAACAGAA TGAAAAAATC	1080
10	GCGCAAATTT ATGCGCTTGA TTTGAAAACT AATGGCGATT ACAACGCTTA TTACAACGCC	1140
	TTGAATGACA AAATCACTAT TCAAATCACC CAGCTTGAAA CCTTAAGCGC TCTAAATAGT	1200
	GCTTATTTGT CCTTACAAAA TCTCAAAGGA TTAGAATGA	1239
	(2) INFORMATION FOR SEQ ID NO:52:	
15		
	(I) SEQUENCE CHARACIERISTICS:	
	(A) LENGTH: 414 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double	
20	(D) TOPOLOGY: circular	
	(ii) MOLECULE TYPE: DNA (genomic)	
0.5	(iii) HYPOTHETICAL: NO	
25		
	(iv) ANTI-SENSE: NO	
	(') OPTGTVVI GOUDGE	
	(vi) ORIGINAL SOURCE:	
30	(A) ORGANISM: Helicobacter pylori	
30	(ix) FEATURE:	
	(A) NAME/KEY: misc_feature	
	(B) LOCATION 1414	
	(D) DOCALLON LLL	
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:52:	
55		
	ATGCGTATAG TTAGAAATTT ATTTCTTGTA TCGTTTGTGG CGTATAGTAG TGCGTTCGCA	60
	GCGGATTTAG AAACCGGAAC CAAAAACGAC AAAAAGAGCG GTAAAAAATT TTACAAACTC	120
	CATAAAAACC ATGGCTCAGA AACCGAGACT AAAAACGATA AAAAGCTTTA TGATTTCACT	180
40	AAAAATAGCG GATTAGAAGG CGTGGATTTA GAAAAAAGCC CTAACCTTAA AAGCCATAAA	240
	AAAAGCGATA AAAAGTTTTA TAAACAACTC GCTAAAAACA ATATCGCTGA AGGGGTGAGC	300
	ATGCCGATTG TGAATTTCAA TAAAGCCCTA TCTTTTGGGC CTTATTTTGA AAGGACTAAA	360
	AGCAAAAAA CCCAATACAT GGACGGCGGG TTGATGATGC ACATCCGTTT TTAA	414
45	(2) INFORMATION FOR SEQ ID NO:53:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 930 base pairs	
	(B) TYPE: nucleic acid	
50	(C) STRANDEDNESS: double	
	(D) TOPOLOGY: circular	
	(ii) MOLECULE TYPE: DNA (genomic)	
	(II) MODECULE TIPE: DNA (Genomic)	

55

(iii) HYPOTHETICAL: NO

	(iv) ANTI-SENSE: NO	
5	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1930</pre>	
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:53:	
	TTGATGCCAC AAAACCAGCT TGTGATCACC ATCATTGATG AATCAGGCTC TAAGCAACTC	60
	AAATTTTCTA AAAATTTAAA ACGCAACCTC ATCATTTCTG TTGTCATTCT TTTATTGATC	120
15	GTGGGGCTTG GCGTGGGGTT TTTAAAATTT TTAATCGCTA AAATGGATAC GATGACAAGC	180
	GAGAGGAATG CGGTTTTAAG GGATTTTAGG GGTTTGTATC AAAAAAATTA CGCCCTAGCG	240
	AAAGAGATTA AAAACAAGCG AGAAGAGCTT TTTATTGTGG GGCAAAAGAT CCGTGGGCTA	300
	GAATCCTTGA TTGAAATCAA AAAGGGGGCT AATGGGGGAG GGCATCTCTA TGATGAAGTG	360
	GATTTAGAAA ATTTGAGCTT AAATCAAAAA CATTTAGCAC TCATGCTCAT TCCTAATGGC	420
20	ATGCCCCTAA AAACTTATAG CGCTATCAAA CCCACTAAAG AAAGGAACCA CCCCATTAAA	480
	AAGATTAAGG GCGTTGAATC CGGGATCGAT TTTATCGCGC CATTGAACAC GCCTGTGTAT	540 600
	GCGAGCGCTG ATGGGATTGT GGATTTTGTG AAGACTCGTT CTAATGCGGG GTATGGGAAC TTGGTGCGCA TTGAACATGC GTTTGGTTTC AGCTCCATTT ATACGCACTT AGATCATGTC	660
	AATGTGCAGC CTAAAAGCTT CATCCAAAAA GGGCAGTTGA TTGGCTATAG CGGGAAGAGC	720
25	GGTAATAGCG GCGGCGAAAA ATTGCATTAT GAAGTGCGGT TTTTGGGTAA AATTTTAGAC	780
23	GCAGAAAAAT TCCTAGCATG GGATTTGGAT CATTTTCAAA GCGCTTTAGA AGAAAATAAA	840
	TTTATTGAAT GGAAGAATCT GTTTTGGGTT TTAGAAGACA TCGTCCAGCT CCAAGAGCAT	900
	GTGGATAAAG ACACCTTAAA AGGTCAGTAG	930
30	(2) INFORMATION FOR SEQ ID NO:54:	
	ALL CONTRACT OVER THE CONTRACT OF THE CONTRACT	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 999 base pairs	
	(B) TYPE: nucleic acid	
35	(C) STRANDEDNESS: double	
55	(D) TOPOLOGY: circular	
	(ii) MOLECULE TYPE: DNA (genomic)	
40	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
	(vi) ORIGINAL SOURCE:	•
45	(A) ORGANISM: Helicobacter pylori	•
	(ix) FEATURE:	
	(A) NAME/KEY: misc_feature	
	(B) LOCATION 1999	
50		
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:54:	
	GTGCTATATT TTTTAACCAG TTTATTTATT TGCTCTTTGA TTGTTTTGTG GTCTAAAAAA	60
	TCCATGCTCT TTGTGGATAA CGCTAATAAA ATCCAAGGCT TCCATCATGC AAGAACCCCA	120
55	CGAGCCGGGG GGCTTGGGAT CTTTCTTTCT TTTGCGTTGG CTTGTTATCT TGAACCTTTT	180

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	GAGATGCCTT TTAAGGGGCC TTTTGTTTTC TTAGGGCTAT CGCTAGTGTT TTTGAGCGGT	240					
	TTTTTAGAAG ACATTAACCT TTCATTAAGC CCCAAAATAC GCCTTATTTT GCAAGCTGTA	300					
	GGGGTCGTTT GCATCATTTC ATCAACGCCT TTAGTGGTGA GCGATTTTTC GCCCCTTTTT	360					
	AGCTTGCCTT ATTTCATCGC TTTTTTATTC GCTATTTTTA TGCTGGTGGG TATCAGTAAC	420					
5	GCTATTAATA TCATTGACGG GTTTAACGGG CTTGCATCTG GGATTTGCGC GATCGCGCTT	480					
	TTAGTCATTC ATTATATAGA CCCTAGCAGT TTGTCTTGTT TGCTCGCTTA CATGGTGCTT	540					
	GGGTTTATGG TGTTAAATTT CCCTTCAGGA AAGATTTTTT TAGGCGATGG GGGGGCGTAT	600					
	TTTTTGGGTT TGGTGTGCGG GATTTCTCTC TTGCATTTGA GTTTGGAGCA AAAAATCAGC	660					
	GTGTTTTTTG GGCTCAATTT AATGCTTTAT CCGGTCATAG AGGTGCTTTT TAGTATCCTT	720					
10	AGGCGCAAAA TAAAACGCCA GAAAGCCACC ATGCCGGATA ATTTGCATTT GCACACCCTT						
10	TTATTTAAAT TCTTGCAACA ACGCTCTTTC AATTACCCTA ACCCTTTATG CGCGTTTATC	780					
		840					
	CTTATTCTAT GCAACCTGCC TTTTATTTTA ATAAGCGTTT TGTTTCGCTT GGACGCTTAT	900					
	GCGCTCATTG TGATTAGCCT AGTCTTTATC GCATGCTATT TAATAGGCTA TGCTTATTTG	960					
1.5	AATAGGCAAG TTTGCGCTTT AGAAAAGCGG GCGTTTTAA	999					
15							
	(2) INFORMATION—FOR SEQ ID NO:55:	-					
	(i) SEQUENCE CHARACTERISTICS:						
	(A) LENGTH: 816 base pairs						
20	(B) TYPE: nucleic acid						
	(C) STRANDEDNESS: double						
	(D) TOPOLOGY: circular						
	(ii) MOLECULE TYPE: DNA (genomic)						
25							
	(iii) HYPOTHETICAL: NO						
	(iv) ANTI-SENSE: NO						
20							
30	(vi) ORIGINAL SOURCE:						
	(A) ORGANISM: Helicobacter pylori						
	44 1						
	(ix) FEATURE:						
2.5	(A) NAME/KEY: misc_feature						
35	·						
35	(A) NAME/KEY: misc_feature (B) LOCATION 1816						
35	(A) NAME/KEY: misc_feature						
35	(A) NAME/KEY: misc_feature (B) LOCATION 1816 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:55:						
	(A) NAME/KEY: misc_feature (B) LOCATION 1816 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:55: ATGAACATAT TCAAGCGTAT TATTTGCGTA ACCGCTATTG TTTTAGGTTT TTTTAACCTT	60					
35 40	(A) NAME/KEY: misc_feature (B) LOCATION 1816 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:55: ATGAACATAT TCAAGCGTAT TATTTGCGTA ACCGCTATTG TTTTAGGTTT TTTTAACCTT TTAGACGCCA AACACCACAA AGAAAAAAAA GAAGACCACA AAATCACTCG TGAGCTTAAA	120					
	(A) NAME/KEY: misc_feature (B) LOCATION 1816 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:55: ATGAACATAT TCAAGCGTAT TATTTGCGTA ACCGCTATTG TTTTAGGTTT TTTTAACCTT TTAGACGCCA AACACCACAA AGAAAAAAAA GAAGACCACA AAATCACTCG TGAGCTTAAA GTGGGCGCTA ACCCTGTGCC GCATGCGCAA ATCTTGCAAT CAGTTGTGGA TGATTTGAAA	120 180					
	(A) NAME/KEY: misc_feature (B) LOCATION 1816 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:55: ATGAACATAT TCAAGCGTAT TATTTGCGTA ACCGCTATTG TTTTAGGTTT TTTTAACCTT TTAGACGCCA AACACCACAA AGAAAAAAA GAAGACCACA AAATCACTCG TGAGCTTAAA GTGGGCGCTA ACCCTGTGCC GCATGCGCAA ATCTTGCAAT CAGTTGTGGA TGATTTGAAA GAGAAAGGGA TCAAATTAGT GATCGTGTCT TTTACGGATT ATGTGTTGCC TAATTTAGCG	120 180 240					
	(A) NAME/KEY: misc_feature (B) LOCATION 1816 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:55: ATGAACATAT TCAAGCGTAT TATTTGCGTA ACCGCTATTG TTTTAGGTTT TTTTAACCTT TTAGACGCCA AACACCACAA AGAAAAAAAA GAAGACCACA AAATCACTCG TGAGCTTAAA GTGGGCGCTA ACCCTGTGCC GCATGCGCAA ATCTTGCAAT CAGTTGTGGA TGATTTGAAA GAGAAAGGGA TCAAATTAGT GATCGTGTCT TTTACGGATT ATGTGTTGCC TAATTTAGCG CTCAATGACG GCTCTTTAGA CGCGAATTAC TTCCAGCACC GCCCTTATTT GGATCGGTTT	120 180 240 300					
40	(A) NAME/KEY: misc_feature (B) LOCATION 1816 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:55: ATGAACATAT TCAAGCGTAT TATTTGCGTA ACCGCTATTG TTTTAGGTTT TTTTAACCTT TTAGACGCCA AACACCACAA AGAAAAAAAA GAAGACCACA AAATCACTCG TGAGCTTAAA GTGGGCGCTA ACCCTGTGCC GCATGCGCAA ATCTTGCAAT CAGTTGTGGA TGATTTGAAA GAGAAAGGGA TCAAATTAGT GATCGTGTCT TTTACGGATT ATGTGTTGCC TAATTTAGCG CTCAATGACG GCTCTTTAGA CGCGAATTAC TTCCAGCACC GCCCTTATTT GGATCGGTTT AATTTGGACA GAAAAATGCA CCTTGTTGGT TTGGCCAATA TCCATGTGGA GCCTTTAAGA	120 180 240 300 360					
	(A) NAME/KEY: misc_feature (B) LOCATION 1816 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:55: ATGAACATAT TCAAGCGTAT TATTTGCGTA ACCGCTATTG TTTTAGGTTT TTTTAACCTT TTAGACGCCA AACACCACAA AGAAAAAAAA GAAGACCACA AAATCACTCG TGAGCTTAAA GTGGGCGCTA ACCCTGTGCC GCATGCGCAA ATCTTGCAAT CAGTTGTGGA TGATTTGAAA GAGAAAGGGA TCAAATTAGT GATCGTGTCT TTTACGGATT ATGTGTTGCC TAATTTAGCG CTCAATGACG GCTCTTTAGA CGCGAATTAC TTCCAGCACC GCCCTTATTT GGATCGGTTT	120 180 240 300 360 420					
40	(A) NAME/KEY: misc_feature (B) LOCATION 1816 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:55: ATGAACATAT TCAAGCGTAT TATTTGCGTA ACCGCTATTG TTTTAGGTTT TTTTAACCTT TTAGACGCCA AACACCACAA AGAAAAAAAA GAAGACCACA AAATCACTCG TGAGCTTAAA GTGGGCGCTA ACCCTGTGCC GCATGCGCAA ATCTTGCAAT CAGTTGTGGA TGATTTGAAA GAGAAAGGGA TCAAATTAGT GATCGTGTCT TTTACGGATT ATGTGTTGCC TAATTTAGCG CTCAATGACG GCTCTTTAGA CGCGAATTAC TTCCAGCACC GCCCTTATTT GGATCGGTTT AATTTGGACA GAAAAATGCA CCTTGTTGGT TTGGCCAATA TCCATGTGGA GCCTTTAAGA TTTTATTCTC AAAAAATCAC AGACATTAAA AACCTTAAAA AAGGCTCAGT GATTGCTGTG CCAAATGATC CGGCCAATCA AGGCAGGGCG TTGATTTTAC TCCATAAACA AGGCCTTATC	120 180 240 300 360 420 480					
40	(A) NAME/KEY: misc_feature (B) LOCATION 1816 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:55: ATGAACATAT TCAAGCGTAT TATTTGCGTA ACCGCTATTG TTTTAGGTTT TTTTAACCTT TTAGACGCCA AACACCACAA AGAAAAAAA GAAGACCACA AAATCACTCG TGAGCTTAAA GTGGGCGCTA ACCCTGTGCC GCATGCGCAA ATCTTGCAAT CAGTTGTGGA TGATTTGAAA GAGAAAGGGA TCAAATTAGT GATCGTGTCT TTTACGGATT ATGTGTTGCC TAATTTAGCG CTCAATGACG GCTCTTTAGA CGCGAATTAC TTCCAGCACC GCCCTTATTT GGATCGGTTT AATTTGGACA GAAAAATGCA CCTTGTTGGT TTGGCCAATA TCCATGTGGA GCCTTTAAGA TTTTATTCTC AAAAAATCAC AGACATTAAA AACCTTAAAA AAGGCTCAGT GATTGCTGTG CCAAATGATC CGGCCAATCA AGGCAGGGCG TTGATTTTAC TCCATAAACA AGGCCTTATC GCTCTCAAAG ACCCCAAGCAA TCTATACGCT ACGGAGTTTG ATATTGTCAA AAATCCCTTAC	120 180 240 300 360 420 480 540					
40	(A) NAME/KEY: misc_feature (B) LOCATION 1816 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:55: ATGAACATAT TCAAGCGTAT TATTTGCGTA ACCGCTATTG TTTTAGGTTT TTTTAACCTT TTAGACGCCA AACACCACAA AGAAAAAAAA GAAGACCACA AAATCACTCG TGAGCTTAAA GTGGGCGCTA ACCCTGTGCC GCATGCGCAA ATCTTGCAAT CAGTTGTGGA TGATTTGAAA GAGAAAGGGA TCAAATTAGT GATCGTGTCT TTTACGGATT ATGTGTTGCC TAATTTAGCG CTCAATGACG GCTCTTTAGA CGCGAATTAC TTCCAGCACC GCCCTTATTT GGATCGGTTT AATTTGGACA GAAAAATGCA CCTTGTTGGT TTGGCCAATA TCCATGTGGA GCCTTTAAGA TTTTATTCTC AAAAAATCAC AGACATTAAA AACCTTAAAA AAGGCTCAGT GATTGCTGTG CCAAATGATC CGGCCAATCA AGGCAGGGCG TTGATTTTAC TCCATAAACA AGGCCTTATC	120 180 240 300 360 420 480 540 600					
40	(A) NAME/KEY: misc_feature (B) LOCATION 1816 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:55: ATGAACATAT TCAAGCGTAT TATTTGCGTA ACCGCTATTG TTTTAGGTTT TTTTAACCTT TTAGACGCCA AACACCACAA AGAAAAAAAA GAAGACCACA AAATCACTCG TGAGCTTAAA GTGGGCGCTA ACCCTGTGCC GCATGCGCAA ATCTTGCAAT CAGTTGTGGA TGATTTGAAA GAGAAAGGGA TCAAATTAGT GATCGTGTCT TTTACGGATT ATGTGTTGCC TAATTTAGCG CTCAATGACG GCTCTTTAGA CGCGAATTAC TTCCAGCACC GCCCTTATTT GGATCGGTTT AATTTGGACA GAAAAATCAC AGACATTAAA AACCTTAAAA AAGGCTCAGT GATTGCTGTG CCAAATGATC CGGCCAATCA AGGCAGGGCG TTGATTTTAC TCCATAAACA AGGCCTTATC GCTCTCAAAG ACCCAAGCAA TCTATACGCT ACGGAGTTTG ATATTGTCAA AAATCCTTAC AACATCAAAA TCAAACCCCT AGAAGCTGCG TTATTGCCTA AGGTTTTAGG GGATGTGGAT GGGGCTATCA TAACAGGGAA TTATGCCTTG CAAGCAAAAC TCACCGGAGC CTTATTTTCA	120 180 240 300 360 420 480 540 600 660					
40	(A) NAME/KEY: misc_feature (B) LOCATION 1816 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:55: ATGAACATAT TCAAGCGTAT TATTTGCGTA ACCGCTATTG TTTTAGGTTT TTTTAACCTT TTAGACGCCA AACACCACAA AGAAAAAAA GAAGACCACA AAATCACTCG TGAGCTTAAA GTGGGCGCTA ACCCTGTGCC GCATGCGCAA ATCTTGCAAT CAGTTGTGGA TGATTTGAAA GAGAAAGGGA TCAAATTAGT GATCGTGTCT TTTACGGATT ATGTGTTGCC TAATTTAGCG CTCAATGACG GCTCTTTAGA CGCGAATTAC TTCCAGCACC GCCCTTATTT GGATCGGTTT AATTTGGACA GAAAAATCAC AGACATTAAA AACCTTAAAA AAGGCTCAGT GATTGCTGTG CCAAATGATC CGGCCAATCA AGGCAGGGCG TTGATTTAC TCCATAAACA AGGCCTTATC GCTCTCAAAG ACCCAAGCAA TCTATACGCT ACGGAGTTTG ATATTGTCAA AAATCCTTAC AACATCAAAA TCAAACCCCT AGAAGCTGCG TTATTGCCTA AGGTTTTAGG GGATGTGGAT	120 180 240 300 360 420 480 540 600					
40	(A) NAME/KEY: misc_feature (B) LOCATION 1816 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:55: ATGAACATAT TCAAGCGTAT TATTTGCGTA ACCGCTATTG TTTTAGGTTT TTTTAACCTT TTAGACGCCA AACACCACAA AGAAAAAAAA GAAGACCACA AAATCACTCG TGAGCTTAAA GTGGGCGCTA ACCCTGTGCC GCATGCGCAA ATCTTGCAAT CAGTTGTGGA TGATTTGAAA GAGAAAGGGA TCAAATTAGT GATCGTGTCT TTTACGGATT ATGTGTTGCC TAATTTAGCG CTCAATGACG GCTCTTTAGA CGCGAATTAC TTCCAGCACC GCCCTTATTT GGATCGGTTT AATTTGGACA GAAAAATCAC AGACATTAAA AACCTTAAAA AAGGCTCAGT GATTGCTGTG CCAAATGATC CGGCCAATCA AGGCAGGGCG TTGATTTTAC TCCATAAACA AGGCCTTATC GCTCTCAAAG ACCCAAGCAA TCTATACGCT ACGGAGTTTG ATATTGTCAA AAATCCTTAC AACATCAAAA TCAAACCCCT AGAAGCTGCG TTATTGCCTA AGGTTTTAGG GGATGTGGAT GGGGCTATCA TAACAGGGAA TTATGCCTTG CAAGCAAAAC TCACCGGAGC CTTATTTTCA	120 180 240 300 360 420 480 540 600 660					
40	(A) NAME/KEY: misc_feature (B) LOCATION 1816 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:55: ATGAACATAT TCAAGCGTAT TATTTGCGTA ACCGCTATTG TTTTAGGTTT TTTTAACCTT TTAGACGCCA AACACCACAA AGAAAAAAA GAAGACCACA AAATCACTCG TGAGCTTAAA GTGGGCGCTA ACCCTGTGCC GCATGCGCAA ATCTTGCAAT CAGTTGTGGA TGATTTGAAA GAGAAAGGGA TCAAATTAGT GATCGTGTCT TTTACGGATT ATGTGTTGCC TAATTTAGCG CTCAATGACG GCTCTTTAGA CGCGAATTAC TTCCAGCACC GCCCTTATTT GGATCGGTTT AATTTGGACA GAAAAATCAC AGACATTAAA AACCTTAAAA AAGGCTCAGT GATTGCTGTG CCAAATGATC CGGCCAATCA AGGCAGGGCG TTGATTTTAC TCCATAAACA AGGCCTTATC GCTCTCAAAG ACCCAAGCAA TCTATACGCT ACGGAGTTTG ATATTGTCAA AAATCCTTAC AACATCAAAA TCAAACCCCT AGAAGCTGCG TTATTGCCTA AGGTTTTAGG GGATGTGGAT GGGGCTATCA TAACAGGGAA TTATGCCTTG CAAGCAAAAC TCACCGGAGC CTTATTTTCA GAAGATAAGG ACTCGCCTTA TGCTAATCTT GTAGCCTCTC GTGAGGATAA TGCGCAAGAT	120 180 240 300 360 420 480 540 600 660 720					

55

(2) INFORMATION FOR SEQ ID NO:56:

5	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 951 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
	(ii)	MOLECULE TYPE: DNA (genomic)	
10	(iii)	HYPOTHETICAL: NO	
10	(iv)	ANTI-SENSE: NO	
	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
15	(ix)	FEATURE: (A) NAME/KEY: misc_feature	
		(B) LOCATION 1951	
20	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:56:	•
	TTAAAGCT	AT TCAGTTTGTG GTGCGATTTT ATAGAAAGGG ATTTTTTAGA AAACGATTTT CA TCAATAAGGG GGCTATTTGC GGGGCGACGA GTAACCCTAG TTTGTTTTGC	60 120
	GAAGCGAT	CA CAAAAAGCGC GTTTTATCAA GATGAAATCG CTAAACTCAA AGGCAAAAAA	180
25		AA TTTATGAAAC TCTGGCACTA AAGGATATTT TACAAGCCTC TAGCGCGTTA	240
		GT ATGAAAAAGA CCCTAACAAC GGCTACATCA GCCTAGAAAT TGACCCCTTT	300
	TTAGAAGA	CG ATGCGATTAA AAGCATTGAT GAAGCCAAGC GGTTATTCAA AACATTAAAC	360
	CGCCCCAA	TG TGATGATTAA AGTCCCGGCG AGTGAAAGCG CTTTTGAAGT CATTAGCGCT	420
	CTGGCTCA	AG CCTCTATCCC CATTAATGTA ACTTTAGTCT TTTCGCCTAA AATTGCCGGT	480 540
30	GAAATCGC	TC AAATCTTAGC CAAAGAAGCA CGAAAAAGAG CGGTCATTAG CGTGTTTGTC	600
	TCACGATT	TG ACAAAGAAT AGACCCACTA GTGCCACAAA ATTTGCAAGC TCAAAGTGGG	660
		CG CTACCGAGTG TTATTATCAA ATCAACCAGC ATGCTAATAA GCTAATAAA	720
	ACCCTTTT	TG CATCCACCGG CGTTAAATCT AATTCTTTAG CTAAAGATTA CTACATTAAA	780
2.5	GCGCTGTG	TT TTAAAAACTC TATCAACACA GCCCCCTAG ACGCCCTAAA CGCTTATTTG	840
35		AA ACACCGAGTG TCAAACCCCT TTAAAAATCA CAGAAATTGA AGCGTTCAAA AA AAACGCACAA TATTGATTTA GAAAACACCG CCCAAAAACT CCTTAAAGAA	900
		AA AAACGCACAA TATIGATITA GAAAACACCG CCCAAAAACT CCTTTTTATATATAA GCAGTTTTTG A	951
	(2) INFO	RMATION FOR SEQ ID NO:57:	
40			
	(i)	SEQUENCE CHARACTERISTICS:	
		(A) LENGTH: 783 base pairs	
		(B) TYPE: nucleic acid	
45		(C) STRANDEDNESS: double (D) TOPOLOGY: circular	
	(ii)	MOLECULE TYPE: DNA (genomic)	
	(iii)	HYPOTHETICAL: NO	
50	(iv)	ANTI-SENSE: NO	
	(vi)	ORIGINAL SOURCE:	
		(A) ORGANISM: Helicobacter pylori	
E E			

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(ix) FEATURE:

	(A) NAME/KEY: misc_feature (B) LOCATION 1783	
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:57:	
•	ATGAAAACAA ATGGTCATTT TAAGGATTTT GCATGGAAAA AATGCTTTTT AGGCGCGAGC	60
	GTGGTGGCTT TATTAGTGGG GTGTAGCCCG CATATTATTG AAACCAATGA AGTTGCTTTG	120
	AAATTGAATT ACCATCCAGC TAGCGAGAAA GTTCAAGCGT TAGATGAAAA GATTTTACTT	180
10	TTAAGGCCAG CTTTCCAATA CAGCGATAAT ATTGCTAAAG AGTATGAAAA CAAATTCAAG	240
	AATCAAACCA CGCTTAAAGT TGAAGAGATC TTGCAAAATC AGGGCTATAA GGTTATTAAT	300
	GTGGATAGCA GCGATAAAGA CGATTTTTCT TTTGCGCAAA AAAAAGAAGG GTATTTGGCT	360
	GTCGCTATGA ATGGCGAAAT TGTTTTACGC CCCGATCCTA AAAGGACCAT ACAGAAAAAA	420
	TCAGAACCCG GGTTATTATT CTCCACTGGT TTGGATAAAA TGGAAAGGGT TTTAATCCCG	480
15	GCTGGGTTTG TCAAGGTTAC CATACTAGAG CCTATGAGTG GGGAATCTTT GGATTCTTTT	540
	ACGATGGATT_TGAGCGAGTT_GGACATCCAA_GAAAAATTCT_TAAAAAGGAG-GGATTCAAGC	-6.00
	CATAGCGGAG GGTTAGTTAG CACTATGGTT AAGGGGACGG ATAATTCTAA TGACGCAATT	660
	AAGAGCGCTT TGAATAAGAT TTTTGCAAGT ATCATGCAAG AAATGGATAA GAAACTCACT	720
	CAAAGGAATT TAGAATCTTA TCAAAAAGAC GCCAAGGAAT TAAAAAACAA GAGAAACCGA	780
20	TAA	783
	(2) INFORMATION FOR SEQ ID NO:58:	
	(i) SEQUENCE CHARACTERISTICS:	
25	(A) LENGTH: 4149 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double	
	(D) TOPOLOGY: circular	
30	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
35	()	
	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Helicobacter pylori	
	(ix) FEATURE:	
40	(A) NAME/KEY: misc feature	
	(B) LOCATION 14149	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:58:	
	(XI) SEQUENCE DESCRIPTION: SEQ ID NO:50:	
45	TTGAATTTTA ATAACCTTAC GGCTAATGGG GCGTTAAATT TTAATGGTTA TGCGCCCTCT	60
	TTAACTAAGG CTTTAATGAA TGTCAGCGGG CAGTTTGTTT TAGGGAATAA TGGGGATATT	120
	AATTTATCTG ACATCAATAT CTTTGACAAC ATCACAAAAT CTGTAACTTA CAACATCTTA	180
	AACGCTCAAA AAGGGATTAC TGGCATTAGT GGGGCTAATG GCTATGAAAA AATCCTTTTT	240
	TATGGCATGA AAATCCAAAA CGCTACCTAT AGCGATAATA ACAACATCCA AACTTGGTCG	300
50	TTTATAAACC CTCTCAATTC TTCTCAAATC ATTCAAGAGA GCATTAAAAA TGGGGATCTA	360
	ACCATAGAAG TTTTAAATAA CCCTAACTCG GCTTCCAACA CTATTTTTAA TATCGCTCCT	420
	GAGCTTTATA ATTACCAAGA TTCTAAGCAA AATCCTACCG GCTATAGCTA TGATTATAGC	480
	GACAATCAAG CAGGCACTTA TTACTTGACA AGCAACATTA AAGGTCTTTT CACCCCTAAA	540
	GGCTCTCAAA CGCCTCAAAC CCCAGGCACT TATAGCCCAT TTAACCAGCC TTTGAATAGT	600

55 TTGAATATCT ACAATAAGGG TTTTTCTAGC GAGAATTTAA AAACGCTTTT AGGGATCCTT 660

	TCTCAAAATT	CCGCCACCTT	AAAAGAAATG	ATTGAATCCA	ACCAACTAGA	CAATATCACT	720
	3 3 C 3 C C C C C C C C C C C C C C C C	AAGTGTTGCA	ACTCTTAGAT	AAGATTAAAA	TCACCCAAGC	GCAAAAGCAA	780
	accompost AC	AAACGATCAA	CCATTTGACT	GACAACATCA	ATCAAACCTT	TAATAACGGG	840
	A RECECCTED	TAGGCGCTAC	CCAAGATAAT	GTTACAAACT	CTACTAGCTC	TATATGGTTT	900
5	CCCCCAATC	GCTATAGCAG	CCCTTGCGCG	CTAGATAGCG	CCACTTGTTC	TTCTTTTAGA	960
5	ላ ላ ር ላ ርጥጥን ርጥ	TEGGGCAATT	ATTAGGCTCA	ACTTCCCCTT	ATTTAGGCTA	CATTAACGCT	1020
	CARROTTAAAC	CTAAAAGCAT	TTATATTACC	GGGACAATTG	GAAGTAGTAA	CGCTTTTGAA	1080
	ACCCACCCA	GCGCGGATGT	AACCTTTCAA	AGCGCTAATA	ACTTAGTGTT	GAATAAAGCT	1140
	AACATAGAAG	CTCAAGCCAC	AGACAATATC	TTTAATCTTT	TGGGTCAAGA	AGGGATTGAT	1200
10	እ እ እ አጥርምምም ነ	ATCAGGGGAA	TTTAGCGAAT	GTTCTTAGTC	AAATGGCTAT	GGAAAAATC	1260
10	AACCAACCCG	GCGGTTTAGG	GAACTTTATA	GAAAACGCTC	TAAGCCCTTT	GAGTAAGGAA	1320
	MARGCAAGCCO	GCTTGCAAGA	TGAAACCTTA	GGCCAACTTA	TAGGTCAAAA	TAACTTAGAT	1380
	G 2 መመመ 2 ጥጥር 2	ATAATAGTGG	AGTCATGAAT	GAAATCCAAA	ACATTATCAG	TCAAAAACTA	1440
	ACCAMMENTE	GCAATTTTGT	TACCCCATCC	ATCATAGAAA	ACTACCTTGC	TAAGCAGTCT	1500
15	AGCAIIIIIG	TGCTAGACGA	TAAAGGGCTT	TTGAATTTTA	TCGGTGGGTA	TATAGACGCT	1560
13	מ מידית א אידית א	CCTCTATTTT	AGGCGTGATT	TTAAAGGATA	TTACTAACCC	CCCTACAAGC	1620
	TCIGAATIAA	ACATTGGTGT	GGTAGCGAAC	GACTTGTTGA	ACGAGTTTTT	AGGACAAGAT	1680
	CIGCMAMA	AGCTAGAAAG	TCAAGGCTTG	GTGAGTAATA	TCATCAATAA	TGTTATTTCT	1740
	CANCECEET	TGAGCGGCGT	TTATAATCAA	GGTTTAGGGA	GCGTGTTGCC	GCCCTCTTTA	1800
20	CAAGGCGGGI	TCAAAGAAAA	CGATTTAGGC	ACTCTTTTAT	CGCCTAGAGG	CTTGCATGAT	1860
20	CAAAACGCGC	AAGGGTATTT	TAACTTTTTA	AGCAATGGCT	ATGTTTTTGT	CAATAACAGC	1920
	TITIGGCAAA	ACGCTACTGG	GGGTAGTTTG	AATTTTGTCG	CCAACAAGTC	TATTATCTTT	1980
	AATCCCCATA	ATACGATTGA	CTTTAGCAAG	TATCAAGGCG	CATTGATTTT	TGCTTCTAAT	2040
	AAIGGCGAIA	ATATCAATAT	CACCACCCTA	AACGCCACTA	ATGGCTTAAG	CCTTAATGCG	2100
25	CCTTTCAATA	ATGTGAGCGT	TCAAAAAGGA	GAAATTTGTA	TCAATTTAGC	CAATTGCCCT	2160
23	GGIIIGAAIA	ACAGCTCTCC	TGCAAACTCT	AGCGTAACCC	CCACTAATGA	GTCTTTAAGC	2220
	CTCCACCCTA	ATAATTTCAC	TTTCTTAGGC	ACAATCATCT	CTAATGGGGC	TATTGATTTG	2280
	GIGCACGCIA GCTCAACTAA	CAAATAATAG	CGTTATAGGC	ACGCTCAATC	TCAATGAAAA	TGCGACCTTG	2340
	መስከርር ምስ አጥል	ATTTAACGAT	CACCAACGCT	TTTAACAACG	CCTCTAACTC	TACGGCTAAT	2400
30	ATTCATCCTA	ATTTCACCTT	AAACCAACAA	GCGACTTTAA	GCACTAACGC	TAGTGGTTTG	2460
30	ATTGATGGTA	GGAATTTTAA	TAGCTATGGC	GATTTGGTGT	TTAACCTCAG	TCATTCAGTT	2520
	ARIGICATO	TTATCAATAC	TCAAGGCACA	GCGACGATCA	TGGCCAATAA	TAACCCTTTG	2580
	AUCCAATTCA	ACGCTTCTTC	AAAAGAAGTG	GGTACTTACA	CGCTGATTGA	TAGCGCTAAA	2640
	CCCATTTATT	ACGGGTATAA	CAACCAAATC	ACAGGAGGCA	GTAGCCTGGA	TAATTACCTT	2700
35	AACCTTTATC	CGCTCATTGA	TATTAATGGC	AAGCACATGG	TGATGACTGA	CAACGGCTTA	2760
33	AAGCITIATG	GGCAAGCCGT	GAGCGTTAAA	GATGGCGGTT	TAGTTGTAGG	CTTTAAGGAC	2820
	MCTCA A A ATC	AATACATTTA	CACTTCCATT	CTTTATAATA	AAGTGAAAAT	CGCTGTTTCT	2880
	AATCATCCTA	TCAATAACCC	ACAAGCCCCC	ACTTTAAAAC	AATATATCGC	TCAAATTCAG	2940
	accommon N N	CCCTCCATAC	CATCGATCAA	GCTGGGGGAA	ATCAAGCGAT	TAATTGGCTC	3000
40	TATE A A A TOT	TTGAAACTAA	AGGAAGCCCT	TTATTCGCTC	CCTATTATCT	AGAGAGCCAC	3060
40	MAINAMAICI	ATTTAACCAC	GATCGCTGGA	GATATTGCTA	ACACTTTAGA	AGTCATCGCT	3120
	A A CCCTTA ATT	TTAAAAATGA	CGCCACTAAT	ATTTTACAGA	TCAACACCTA	CACGCAGCAA	3180
	ARCACTCATT	TAGCCAAGCT	CTCTGACACT	TCAACTTTCG	CCCGTTCTGA	TTTCTTAGAA	3240
	AIGAGICGII	CCCTTAAAAA	CAAGCGATTC	GCTGATGCGA	TCCCTAACGC	TATGGATGTG	3300
15	A COCTIAGAAG	ACTCTCAAAG	GAATAGAGTT	AAAAATAATG	TGTGGGCGAC	AGGAGTTGGA	3360
45	ATTITAAAAT	TCATTAGTGG	AGGTACTGGA	ACTTTATATG	GTATCAATGT	AGGGTATGAT	3420
	GGGGCTAGTI	ACCCCCTGAT	TGTGGGAGGT	TATGCCGCTT	ATGGGTATAG	CGGGTTCCAT	3480
	AGGITTATTA	CTCDATCACC	CTCTAGCAAT	GTCAATGTGG	GCGTTTATAG	CCGAGCGTTT	3540
	GCAAACATCA	CICARICAGO CCCACCTAAC	CATGAGCTTG	AATGAGACTT	GGGGATACA	TAAAACTTTC	3600
60	ATCAAAAGAA	, ALCAGCIANC	ACTCTCAATC	ATCAATCAGT	CTTACAGATA	CGACACTTGG	3660
50	ATCAACTCCT	. CHNNYNMCAN	TTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTT	GATTTCATCT	TTAAAGATA	AAGCGTTATT	3720
	ACGACTGACG	· YYGMYGGGGG	י אסטטיייאיי	TACATTGGT	TGTCTGGTT	AAGGGGCATT	3780
	TTTAAACCCC	, AAGIAGGCII	. אאפרואוואו	GCCAATGCTC	ACCCTAATA	AAAATCCGTT	3840
	ATGGATGATC	. CIMILIACAA	ACD DACTOCO	CATTATTTC	ATAAAAACTO	TTATTATTTT	3900
ے ہے	CTAACGATCA	ATTITUCCO	. אכאראט ז כטכ י אכארידיאידיר	ATTAATTCT	TGGGGGATA	AATGGTGCGT	3960
55	GTGATTGCGG	WIGIGGCWG					

e	TTCATCGGTA ATAACACCCT AAGCTATAGA GATGGTGGCA GATACAACAC TTTTGCTAGC ATTATCACAG GCGGGAGAT AAGATTGTTC AAAACCTTTT ATGTGAATGC GGGCATAGGG GCTAGGTTTG GGCTTGATTA TAAAGATATT AATATTACCG GAAATATTGG TATGCGCTAT GCTTTTTAA	4020 4080 4140 4149
. 5	(2) INFORMATION FOR SEQ ID NO:59:	
10	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 789 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
15	(ii) MOLECULE TYPE: DNA (genomic)	
-	(iii) HYPOTHETICAL: NO.	-
	(iv) ANTI-SENSE: NO	
20	(vi) ORIGINAL SOURCE:(A) ORGANISM: Helicobacter pylori	
25	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1789</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:59:	
30	ATGAAAAAA TTGGTTTGAG CTTGTGTTTG GTTTTGAGTT TGGGTTTTTT AAAAGCCCAT GAAGTGAGCG CTGAAGAGAT TGCGGATATT TTCTACAAAC TCAACGCCAA AGAGCCTAAA ATGAAAATCA ACCACACGAA GGGGTTTTGC GCTAAAGGCG TGTTCCTCCC TAACCCGCAA GCAAGAGAGAG ATTTAGAGGT GCCACTACTC AATGAAAAAG AAATCCCTGC GTCTGTAAGG TATTCTTTAG GGGGCGTGGC GATGGACGAT AAAAGCAAGG TTAGGGGAAT GGCGTTAAAA	60 120 180 240
35	CTAGAAAATC AAAACGCTAG TTGGACAATG GTGATGCTCA ATACAGAAAT CAATTTTGCC AAAAACCCTG AAGAATTCGC CCAATTTTTT GAAATGAGAC TTCCTAAAAA TGGCAAGGTA GATGAAGCAA GAATCAAAAA GCTTTACGAA GAAGTCCCCT CTTATAGGAA TTTTGCCGCC TATATGAAAA CGATAGGGAT TAGCTCAAGC GTGGCTAATA CGCCTTATTA TAGCGTGCAT GCGTTCAAGT TTAAAGATAA GAAAGAAAAA TTATTGCCTG CGAGGTGGAA ATTTGTGCCT	300 360 420 480 540 600
40	AAAGAGGGCG TTAAATACTT AAATCCTCAA GAATTAAAGC AAAAAGATTC AAATTATCTG CTCTCTTCAT TCCAACAACA CCTTAAAAAT AAACCCATAG AATACCAAAT GTATTTGGTG TTTGCGAATC AAAATGATGC CACCAACGAC ACGACCGCGC TTTGGAAAGG CAGCATAAGG AATTATTAG	660 720 780 789
45	(2) INFORMATION FOR SEQ ID NO:60:	
50	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 741 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	

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	(iv) ANTI-SENSE: NO	
~	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
5	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1741</pre>	
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:60:	
15	ATGAAACAAT TTAAAAAGAA ACCAAAAAAAG ATAAAACGAT CGCATCAAAA TCAAAAAAACA ATCTTAAAGC GTCCTTTATG GCTTATGCCT TTACTGATTG GCGGGTTTGC TAGTGGGGTG TATGCGGATG GAACAGACAT TTTGGGGCTT AGTTGGGGG AAAAAAGCCA AAAGGTATGC GTGCATCGTC CATGGTATGC TATATGGAGT TGCGATAAAT GGGAGGAAAA AACACAACAA TTTACAGGAA ACCAACTCAT CACAAAAACT TGGGCAGGGG GTAATGCGGC TAACTACTAC	60 120 180 240 300
	CACTCTCAAA ACAACCAAGA CATCACAGCC AATTTAAAAA ATGATAACGG CACTTATTTT TTAAGCGGTC TGTATAACTA CACCGGAGGG GAATATAATG GGGGGAATTT AGACATTGAA TTAGGCAGTA ACGCTACTTT TAATCTAGGT GCGAGTAGTG GGAATAGCTT CACTTCTTGG	360 420 480
20	TATCCTAATG GGCATACTGA TGTTACTTTT AGCGCTGGGA CTATCAATGT GAATAACAGC GTAGAAGTGG GCAATCGTGT GGGATCGGGA GCTGGCACG ACACCGGCAC AGCCACTTTA AACTTGAACG CTAATAAGGT TACTATCAAT TCCAATATCA GCGCGTATAA AACTTCGCAA GTGAATGTAG GCAATGCTAA CAGCGTTATT ACCATTAATT CGGTTTCTTT AAATGGGGAA	540 600 660 720 741
25	TACTTGCAGT TCTTTAGCTA G (2) INFORMATION FOR SEQ ID NO:61:	,
30	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 738 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
35	(ii) MOLECULE TYPE: DNA (genomic) (iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
40	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
45	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1738</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:61:	
50	ATGATAAAA AGACCCTTGC ATCGGTTTTA TTAGGATTGA GTTTGATGAG TGTGTTAAAT GCCAAAGAAT GCGTTTCGCC CATAACAAGA AGCGTTAAGT ATCATCAGCA AAGTGCTGAG ATCAGAGCCT TGCAATTACA AAGTTACAAA ATGGCGAAAA TGGCGCTAGA CAATAACCTT AAGCTCGTTA AAGACAAAAA GCCAGCCGTC ATCTTGGATT TAGATGAAAC CGTTTTGAAC ACTTTTGATT ATGCGGGCTA TTTAGTCAAA AACTGCATTA AATACACCCC AGAAACTTGGATAAATTTG AAAAAGAAGG CTCTCTTACG CTCATTCCTG GAGCGCTAGA CTTTTTAGAA	120 180 240 300 360
55	TACGCTAATT CTAAGGGCGT TAAGATTTTT TACATTTCTA ACCGCACCCA AAAAAATAAG	420

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	GCATTCACTT TAAAAACGCT CAAAAGCTTT AAGCTCCCCC AAGTGAGTGA AGAATCCGTT TTGTTAAAGG AAAAAGGCAA GCCTAAAGCC GTTAGGCGGG AGTTAGTCGC TAAGGATTAT GCGATTGTTT TACAAGTGGG CGACACTTTG CATGATTTTG ACGCCATTTT TGCTAAAGAC	480 540 600
	GCTAAAAACA GCCAAGAACA ACAAGCCAAA GTCTTGCAAA ACGCTCAAAA ATTCGGCACA	660
5	GAATGGATCA TTTTACCCAA CTCTCTTTAT GGCACATGGG AAGATGGGCC TATAAAAGCA	720
	TGGCAAAATA AAAAATAA	738
	(2) INFORMATION FOR SEQ ID NO:62:	
10	(i) SEOUENCE CHARACTERISTICS:	
10	(A) LENGTH: 867 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double	
	(D) TOPOLOGY: circular	
15		
	_(ii)_MOLECULE -TYPE:DNA(genomic)	-
	(111) INCOMPRESSOR AND	
	(iii) HYPOTHETICAL: NO	
20	(iv) ANTI-SENSE: NO	
	(=1, ===================================	
	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Helicobacter pylori	
25		
25	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature</pre>	
	(B) LOCATION 1867	
	(2) 200:1201 211100	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:62:	
30		
	TTGTGGTGTT TAAAAACCCC TATCATAGGG CATGGCATGA AGAAAAAAGC AAAAGTCTTT	60
	TGGTGTTGTT TTAAAATGAT TCGTTGGTTG TATTTGGCGG TCTTTTTTT GTTGAGCGTA	120
	TCAGACGCTA AAGAAATCGC TATGCAACGA TTTGACAAAC AAAACCATAA GATTTTTGAA ATCCTTGCGG ATAAAGTGAG CGCCAAAGAC AATGTGATAA CCGCCTCAGG GAATGCGATC	180 240
35	CTATTGAATT ATGACGTGTA TATTCTAGCG GATAAGGTGC GTTATGACAC CAAGACTAAA	300
55	GAAGCGTTAT TAGAAGGCAA TATTAAGGTT TATAGGGGCG AGGGCTTGCT CGTTAAAACC	360
	GATTATGTGA AATTGAGTTT GAACGAAAAA TATGAGATCA TTTTCCCCTT TTATGTCCAA	420
	GACAGCGTGA GCGGGATTTG GGTGAGCGCG GATATTGCTA GCGGGAAGGA TCAAAAATAT	480
	AAGATTAAAA ACATGAGCGC TTCAGGGTGC AGCATTGACA ACCCCATTTG GCATGTCAAT	540
40	GCGACTTCAG GCTCATTTAA CATGCAAAAA TCGCATTTGT CAATGTGGAA TCCTAAGATT	600
	TATGTCGGCG ATATTCCTGT ATTGTATTTG CCCTATATTT TCATGTCCAC GAGCAATAAA	660
	AGAACTACCG GGTTTTTATA CCCTGAGTTT GGCACTTCCA ACTTAGACGG CTTTATTTAT	720
	TTGCAACCCT TTTATTTAGC CCCCAAAAAC TCATGGGATA TGACCTTTAC CCCACAAATC CGTTACAAAA GGGGTTTTGG CTTGAATTTT GAAGCGCGCT ACATCAACTC TAAGACGCAG	780 840
45	GTTTTTATTC AATGCGCGCT ATTTTAG	867
15	UIIIIAIIC AAIGCCCCI AIIIIAG	
	(2) INFORMATION FOR SEQ ID NO:63:	
	•	
0.	(i) SEQUENCE CHARACTERISTICS:	
50	(A) LENGTH: 387 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double	
	(D) TOPOLOGY: circular	
55	(ii) MOLECULE TYPE: DNA (genomic)	
~ -	·	

	(iii) HYPOTHETICAL: NO	
_	(iv) ANTI-SENSE: NO	
5	(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
10	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1387</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:63:	
15	TTGATGTTTA AAAAAATGTG TTTGAGCCTG CTAATGATAA GCGGTGTTTG TGTGGGGGCA AAGGATTTGG ATTTCAAGCT GGATTATCGC GCGACTGGGG GGAAATTCAT GGGGAAAATG ACGGACTCTA GTCTTTTAAG TATCACTTCT ATGAACGATG AACCGGTGGT GATTAAAAAC CTTATTGTCA ATAGGGGAAA TTCATGCGAA GCGACTAAAA AAGTAGAACC CAAATTTGGC	60 120 180 240
20	GATAAGTTTA AAAAAGAAAA ACTCTTTGAT CATGAATTAA AATACTCGCA ACAGATATTT TACCGCCTGG ATTGCAAGCC TAACCAATTG TTAGAAGTTA AAATCATCAC GGACAAGGGC GAATATTACC ATAAATTTTC CAAATAG	300 360 387
	(2) INFORMATION FOR SEQ ID NO:64:	
25	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 510 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double	
30	(D) TOPOLOGY: circular	
	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
35	(iv) ANTI-SENSE: NO	
	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
40	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1510</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:64:	
45	ATGCAAGCGT TAAAATCATT GCTTGAAGTG ATTACAAAAC TCCAGAATCT AGGCGGCTAT TTGATGCATA TAGCTATTTT CATCATTTTT ATTTGGATTG GAGGGCTTAA GTTTGTGCCT TACGAAGCTG AAGGGATCGC CCCTTTTGTG GCCAACTCCC CTTTCTTTC TTTCATGTAT AAATTTGAAA AACCTGCATA CAAACAACAC AAAATGTCTG AATCCCAATC CATGCAAGAA	60 120 180 240
50 55	GAAATGCAAG ATAACCCTAA AATCGTTGAA AACAAAGAAT GGCATAAAGA AAACCGCACT TATTTAGTGG CTGAAGGTTT AGGGATTACG ATCATGATCC TAGGCATTTT GGTGCTTTTG GGGCCTTTGGA TGCCTTTAAT GGGCGTAGTT GGGGGCTTGC TTGTCGCTGG AATGACGATC ACCACCCTAT TCTTTTTTAT TCACAACGCC AGAAGTGTTT GTCAATCAGC ATTTCCCATG GCTTTCTGGG GCTGGAAGGC TAGTGGTTAA	300 360 420 480 510

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(2) INFORMATION FOR SEQ ID NO:65:

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(i) SEQUENCE CHARACTERISTICS:
               (A) LENGTH: 1464 base pairs
 5
               (B) TYPE: nucleic acid
               (C) STRANDEDNESS: double
               (D) TOPOLOGY: circular
         (ii) MOLECULE TYPE: DNA (genomic)
10
        (iii) HYPOTHETICAL: NO
        (iv) ANTI-SENSE: NO
15
         (vi) ORIGINAL SOURCE:
       -(A)-ORGANISM:-Helicobacter-pylori
         (ix) FEATURE:
               (A) NAME/KEY: misc_feature
20
               (B) LOCATION 1...1464
         (xi) SEQUENCE DESCRIPTION: SEQ ID NO:65:
     ATGATTGAAT GGATGCAAAA TCATAGAAAG TATTTAGTGG TTACGATATG GATAAGCACG
     ATCGCTTTTA TTGCCGCCGG AATGATAGGT TGGGGGCAAT ACAGCTTTTC TTTAGATAGC
25
     GATAGCGCTG CCAAAGTGGG ACAGATTAAG ATTTCTCAAG AAGAATTAGC CCAAGAATAC
                                                                        180
     CGCCGCCTTA AAGACGCCTA TGCTGAGTCT ATCCCTGATT TTAAAGAACT CACCGAAGAT
     CAAATCAAAG CCATGCATTT AGAAAAAAGC GCGCTAGATT CGCTCATCAA TCAAGCTTTA
     TTGAGGAATT TCGCTTTAGA TTTAGGGCTT GGTGCTACCA AGCAAGAAGT GGCCAAAGAG
     ATCAGAAAAA CGAACGTTTT TCAAAAAGAT GGCGTTTTTG ATGAAGAATT GTATAAAAAT
                                                                        420
30
     ATCTTAAAAC AAAGCCATTA CCGCCCCAAG CATTTTGAAG AAAGCGTTGA AAGGCTTTTA
                                                                        480
     ATCCTTCAAA AAATCAGCGC TCTATTCCCC AAAACCACCA CCCCTTTGGA GCAATCCAGT
                                                                        540
     CTATCGCTTT GGGCAAAATT GCAAGACAAA TTAGACATTC TTATCCTAAA TCCTAATGAT
                                                                        600
     GTTAAAATCT CTCTCAATGA AGAAGAGATG AAAAAATATT ATGAAAACCA TAGAAAGGAT
                                                                        660
35
     TTTAAAAAGC CCACAAGCTT TAAAACACGC TCTTTATATT TTGACGCTAG TTTAGAAAAA
                                                                        720
                                                                        780
     ACTGATTTGA AAGAGTTGGA GGAATACTAC CATAAAAACA AGGTGTCTTA TTTGGACAAA
     GAGGGGAAAT TACAGGATTT TAAAAGCGTT CAAGAGCAAG TCAAGCATGA TTTAAACATG
     CAAAAGGCGA ATGAAAAAGC CTTAAGGAGC TATATCGCTC TAAAAAAGGG GAACGCACAA 900
     AACTACACCA CGCAAGATTT TGAAAAAAAC AACTCCCCCT ATACTGCTGA AATCACGCAA
                                                                        960
     AAACTCACCG CTCTCAAGCC CCTTGAAGTC CTAAAACCAG AGCCTTTTAA AGATGGTTTT 1020
40
     ATCGTGGTGC AGCTTGTCTC TCAAATTAAA GACGAATTGC AAAATTTTGA TGAAGCCAAA 1080
     AGCGCTCTTA AAACCCGTCT GACTCAAGAA AAAACCCTTA TGGCGTTGCA AACTTTAGCT 1140
     AAAGAAAGC TTAAGGATTT TAAAGGGAAA AGCGTGGGTT ATGTAAGCCC TAATTTTGGA 1200
    GGCACTATCA GTGAACTTAA CCAAGAAGAG AGCGCGAAGT TTATCAACAC CCTTTTTAAC 1260
45
     CGCCAGGAAA AAAAAGGGTT TGTAACCATA GGTAATAAAG TGGTGCTTTA TCAAATCACA 1320
     GAGCAAAATT TCAATCACCC CTTTAGTGCA GAAGAAAACC AATACATGCA GCGTTTAGTC 1380
     AATAACACTA AAACGGATTT TTTTGATAAA GCGTTGATAG AAGAATTGAA AAAACGCTAT 1440
                                                                       1464
    AAGATAGTCA AATACATTCA ATAA
50
    (2) INFORMATION FOR SEQ ID NO:66:
          (i) SEQUENCE CHARACTERISTICS:
               (A) LENGTH: 429 base pairs
              (B) TYPE: nucleic acid
55
              (C) STRANDEDNESS: double
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	(D) TOPOLOGY: circular	
	(ii) MOLECULE TYPE: DNA (genomic)	
5	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
10	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1429</pre>	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:66:	
20	ATGAAAACGA ACTTTATAA AATTAAATTA CTATTGCTT GGTGTCTTAT CATTGGCATG TTTAACGCTC CGCTTAACGC TGACCAAAAC ACGGATATAA AAGATATTAG TCCTGAAGAT ATGGCGCTAA ATAGCGTGGG GCTTGTTTCT AGAGATCAGC TAAAAATAGA GATCCCTAAA GAAACCCTAG AGCAAAAAGT GGCCATACTC AATGACTATA ATGATAAGAA TGTTAATATC AAGTTTGACG ACATAAGTTT AGGGAGTTTC CAACCTAATG ATAATCTAGG TATCAATGCG ATGTGGGGCA TTCAAAATCT TCTCATGAGC CAAATGATGA GCAATTACGG TCCAAACAAT	60 120 180 240 300 360
25	TCTTTCATGT ATGGCTATGC GCCAACATAC TCAGATTCAT CGTTTTTACC ACCGATCTTA GGGTATTAA	420 429
	(2) INFORMATION FOR SEQ ID NO:67:	
30	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 627 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
35	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
40	(iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
	(ix) FEATURE:	
45	(A) NAME/KEY: misc_feature (B) LOCATION 1627	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:67:	
50	TTGATCAACA ATAATAATAA CAATAAAAAA CTGAGAGGCT TTTTTTTGAA AGTTCTCTTA AGTCTCGTTG TTTTCAGTTC GTATGGGTCA GCAAATGACG ATAAAGAAGC CAAAAAAGAA GCGCTAGAAA AAGAAAAAAA CACTCCCAAT GGGCTTGTTT ATACGAATTT AGATTTTGAT AGTTTTAAAG CGACTATCAA AAATTTGAAA GACAAGAAAG TAACTTTCAA AGAAGTCAAT	120 180 240 300
55	CCCGATATTA TCAAAGATGA AGTTTTTGAC TTCGTGATTG TCAATAGAGT CCTTAAAAAA ATAAAGGATT TGAAGCATTA CGATCCAGTT ATTGAAAAAA TCTTTGATGA AAAGGGTAAA	360

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	GAAATGGGAT TGAATGTAGA ATTACAGATC AATCCTGAAG TGAAAGACTT TTTTACTTTC	420
	AAAAGCATCA GCACGACCAA CAAACAACGC TGCTTTCTAT CATTGCACGG AGAAACAAGA	480
	GAAATTTTAT GCGATGATAA GCTATATAAT GTTTTATTGG CCGTATTCAA TTCTTATGAT	540
	CCTAATGATC TTTTGAAACA CATTAGCACC ATAGAGTCTC TCAAAAAAAT CTTTTATACG	600
5	ATTACATGTG AAGCGGTATA TCTATAA	627
5	ATTACATOR ANGCOGRAFA TOTALLA	5 _,
	(2) INFORMATION FOR SEQ ID NO:68:	
	(i) SEQUENCE CHARACTERISTICS:	
10	(A) LENGTH: 738 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double	
	(D) TOPOLOGY: circular	
15	(ii) MOLECULE TYPE: DNA (genomic)	
-((1111) INCOMPRESCRIP AND	
	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
20	(IV) ANII-SENSE. NO	
20	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Helicobacter pylori	
	(iii) Olici Zilaiiii iiidaa aa Fii aa	
	(ix) FEATURE:	
25	(A) NAME/KEY: misc feature	
	(B) LOCATION 1738	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:68:	
30	ATGGCAGGCA CACAAGCTAT ATATGAATCA TCTTCTGCAG GATTCTTATC GCAAGTCTCC	60
30	TCAATCATCT CAAGCACAAG TGGTGTCGCA GGGCCATTTG CAGGAATAGT AGCGGGCGCT	120
	ATGACAGCAG CGATTATTCC TATTGTTGTG GGATTTACTA ATCCGCAAAT GACCGCTATC	180
	ATGACCCAAT ACAATCAAAG CATCGCTGAA GCTGTAAGCG TGCCTATGAA AGCCGCTAAC	240
	CAACAATACA ACCAATTGTA TCAAGGTTTT AACGATCAAA GCATGGCTGT GGGGAACAAT	300
35	ATCTTAAATA TCAGCAAATT AACAGGGGAA TTTAACGCGC AAGGCAACAC GCAAAGCGCG	360
33	CAAATTAGTG CTGTCAATAG TCAGATTGCA AGCATTTTAG CGAGTAACAC TACCCCTAAA	420
	AATCCTAGCG CTATTGAAGC TTATGCGACG AATCAAATCG CTGTTCCTAG CGTGCCAACA	480
	ACGGTTGAAA TGATGAGCGG TATATTAGGC AATATTACAA GCGCAGCACC AAAATACGCC	540
	CTAGCTCTAC AAGAGCAACT GCGTTCTCAA GCAAGCAACA GCTCAATGAA TGATACAGCC	600
40	GATTCCCTTG ATAGCTGTAC CGCTTTAGGC GCACTTGTTG GCTCATCAAA AGTGTTTTTC	660
	AGTTGCATGC AAATTTCTAT GACTCCTATG AGTGTTTCTA TGCCCACTGT TATGCCAAAT	720
	ACCAGCGGTT GCCACTAA	738
	(2) INFORMATION FOR SEQ ID NO:69:	
45		
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 1104 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double	
50	(D) TOPOLOGY: circular	
	(ii) MOLECULE TYPE: DNA (genomic)	

55

(iii) HYPOTHETICAL: NO

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(iv) ANTI-SENSE: NO
         (vi) ORIGINAL SOURCE:
               (A) ORGANISM: Helicobacter pylori
5
         (ix) FEATURE:
               (A) NAME/KEY: misc_feature
               (B) LOCATION 1...1104
         (xi) SEQUENCE DESCRIPTION: SEQ ID NO:69:
10
    ATGATTAAAA GCGTAGAGAT TGAAAATTAC AAAAATTTTG AGCACCTTAA AATGGAAAAT
     TTTAAACTCA TCAACTTTTT TACCGGTCAA AACGATGCGG GTAAAACCAA TCTTTTAGAA
     GCTCTTTATA CCAACACAGG CCTTTGTGAT CCTACTGCCA ATCAAGTCAG TCTTCCTCCT
    GAACATGCCG TGAATATTAG TGAATTCAGA AAAATCAAAC TCGATGCCGA CAACCTAAAA
15
     ACCTTTTTT ATCAAGGAAA CACCGCTAAT CCCATTAGTA TCCGCACTGA ATTTGAACAT
     GCTACTATCC CTCTTACTAT CCAATACCCC ACACAAACCA GTTACAGCAA AGACATCAAT
                                                                          360
     TTGAATAGCG ATGATGCTCA TATGACAAAC CTTATAAACA CAACAATAAC GAAGCCACAG
                                                                          420
     CTCCAATTT CCTACAATCC ATCCCTTTCC CCCATGACAA TGACTTATGA ATTTGAAAGG
                                                                          480
     CAAAACCTAG GTTTAATCCA TTCTAATTTA GATAAAATCG CTCAAACCTA TAAAGAAAAT
                                                                          540
20
     GCGATGTTTA TTCCTATAGA ATTATCTATT GTTAATTCTC TTAAAGCATT GGAAAATTTA
                                                                          600
     CAATTAGCAA GCAAAGAAAA AGAATTGATT GAAATCCTAC AATGTTTCAA CCCTAATATT
                                                                          720
     TTAAATGCTA ATACAATAAG AAAGTCTGTC TATATCCAAA TCAAAGATGA AAACACACCG
                                                                          780
     CTAGAAGAAA GTCCCAAAAG GCTTTTAAAT TTGTTTGGTT GGGGTTTTAT CAAATTCTTT
     ATTATGGTGA GCATTCTTAT AGACAATCGT GTCAAGTATC TTTTTATTGA TGAAATAGAA
                                                                          840
25
     AGCGGTTTGC ACCATACAAA AATGCAAGAG TTTTTAAAAG CTCTGTTTAA GTTAGCTCAA
                                                                          900
     AAATTACAGA TTCAAATTTT TGCCACCACG CACAATAAGG AATTTTTATT AAACGCCATC
     AACACGATAT CCGATAATGA AACGGGAGTT TTTAAAGACA TAGCCTTGTT TGAGCTTGAA 1020
     AAAGAAAGCG CTTCTGGCTT TATCAGACAC AGCTATTCTA TGCTAGAAAA AGCGCTTTAT
                                                                         1080
                                                                         1104
     AGGGGTATGG AGGTTAGAGG CTGA
30
     (2) INFORMATION FOR SEQ ID NO:70:
          (i) SEQUENCE CHARACTERISTICS:
               (A) LENGTH: 1230 base pairs
35
               (B) TYPE: nucleic acid
               (C) STRANDEDNESS: double
                (D) TOPOLOGY: circular
        (ii) MOLECULE TYPE: DNA (genomic)
40
        (iii) HYPOTHETICAL: NO
         (iv) ANTI-SENSE: NO
45
         (vi) ORIGINAL SOURCE:
               (A) ORGANISM: Helicobacter pylori
         (ix) FEATURE:
50
                (A) NAME/KEY: misc feature
                (B) LOCATION 1...1230
          (xi) SEQUENCE DESCRIPTION: SEQ ID NO:70:
     ATGTCCTTGA TTAGAGTGAA TGGGGAAGCT TTTAAACTCT CTTTGGAAAG TTTAGAAGAA
55
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	GATCCTTTTG AAACTAAAGA AACGCTAGAA ACGCTAGAAA CGCTTATCAA ACAAACGAGC	120
	GTTGTTTTAT TGGCCGCTGG GGAGTCTAAG CGTTTTTCTC GTGCGATTAA AAAGCAGTGG	180
	CTACGCTCTC ACCACACCCC CTTATGGCTC AGCGTGTATG AAAGCTTTAA AGAAGCCCTA	240
	GACTTTAAGG AAGTCATTCT AGTTGTAAGC GAATTGGATT ATGTTTATAT CCAACGCCAT	
5	TACCCCAAAA TCAAGCTTGT AAAAGGCGGG GCATCAAGGC AAGAATCCGT GCGTAACGCT	360
•	TTGAAAGTAA TTGATAGCAC TTACACGATC ACCAGCGATG TGGCTAGGGG TTTAGCGAAT	420
	ATGGAAGCGC TTAAAAGCTT GTTTTTAACC CTCCAACAAA CGAGCCATTA TTGCATCGCC	480
	CCTTACTTGC CTTGCTATGA CACAGCGATC TATTATAACG AGGCTTTAGA TAGAGAAGCG	540
	ATCAAACTCA TTCAAACCCC GCAATTAAGC CACACCAAAA CGCTCCAATC AGCCCTAAAC	600
10	CAAGGGGGTT TTAAAGATGA AAGCAGCGCG ATTTTACAAG CTTTCCCTAA CTCTGTGAGC	660
	TATATTGAAG GCAGTAAGGA TTTGCACAAA CTCACCACAA GCGGCGATTT AAAGTTTTTT	720
	ACGCCTTTTT TTAACCCAGC AAAGGACACT TTTATAGGCA TGGGTTTTGA TACGCATGCG	780
	TTCATTAAAG ATAAGCCTAT GGTTTTAGGG GGGGTTGTTT TGGATTGCGA GTTTGGGTTA	840
1.6	AAGGCTCATA GCGATGGCGA TGCTTTATTG CATGCGGTTA TTGATGCGAT TTTAGGAGCG	900
15	ATTAAAGGGG GGGATATTGG CGAATGGTTC CCTGATAATG ACCCCAAATA CAAAAACGCC	960
	TETTETAAAG AGETTTTAAA AATCGTGTTG GATTTTCTC AAAGCATTGG GTTTGAATTG	1020
	CTTGAAATGG GAGCGACCAT CTTTAGCGAA ATCCCTAAAA TCACTCCTTA CAAACCGGCG	1080
	ATTTTAGAGA ATTTGAGCCA ACTTTTGGGT TTAGAAAAAT CTCAAATCAG CTTGAAAGCC ACTACAATGG AAAAAAATGGG GTTCATTGGC AAACAAGAAG GGCTGTTAGT CCAAGCGCAT	1140
20	GTGAGCATGC GTTATAAACA AAAACTTTAA	1200
20	GIGAGCAIGC GITATAAACA AAAACIITAA	1230
	(2) INFORMATION FOR SEQ ID NO:71:	
	(2) 2.11 0.12 1.01 0.12 1.01 1.12	
	(i) SEQUENCE CHARACTERISTICS:	
25	(A) LENGTH: 813 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double	
	(D) TOPOLOGY: circular	
30	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
26	(iv) ANTI-SENSE: NO	
35		
	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Helicobacter pylori	
40	(ix) FEATURE:	
40	(A) NAME/KEY: misc_feature	
	(B) LOCATION 1813	
	(xi) SEQUENCE DESCRIPTION: SEO ID NO:71:	
•	(XI) SEQUENCE DESCRIPTION: SEQ ID NO:/I:	
45	ATGAAAAGT TTGTAGCTTT AGGGCTTCTA TCCGCGGTTT TAAGCTCTTC GTTGTTAGCC	60
,,,	GAAGGTGATG GTGTTTATAT AGGGACTAAT TATCAGCTTG GACAAGCCCG TTTGAATAGC	120
	AATATTTATA ATACAGGGGA TTGCACAGGG AGTGTTGTAG GTTGCCCCCC AGGTCTTACC	180
	GCTAATAAGC ATAATCCAGG AGGCACCAAT ATCAATTGGC ACTCCAAATA CGCTAATGGG	240
	GCTTTGAATG GTTTTGGGTT GAATGTGGGT TATAAGAAAT TCTTCCAATT CAAGTCGCTA	300
50	GATATGACAA GCAAGTGGTT TGGTTTTAGA GTGTATGGGC TTTTTGATTA CGGGCATGCC	360 360
20	GATATGACAA GCAAGTGGTT TGGTTTTAGA GTGTATGGGC TTTTTGATTA CGGGCATGCC GATTTAGGTA AACAAGTTTA TGCACCTAAT AAAATCCAGT TGGATATGGT CTCTTGGGGT	420
	GTGGGGAGCG ATTTGTTAGC TGATATTATT GATAAAGACA ACGCTTCTTT TGGTATTTTT	420
	GGTGGGGTCG CTATCGGCGG TAACACTTGG AAAAGCTCTG CAGCAAACTA TTGGAAAGAG	540
	CAAATCATTG AAGCCAAAGG TCCTGATGTT TGTACCCCTA CTTATTGTAA CCCTAATGCC	600
55	CCTTATAGCA CCAACACTTC AACCGTCGCT TTTCAAGTGT GGTTGAATTT TGGGGTGAGA	660

CCTTATAGCA CCAACACTTC AACCGTCGCT TTTCAAGTGT GGTTGAATTT TGGGGTGAGA

660

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	GCCAATATCT ACAAGCATAA TGGCGTGGAA TTTGGCGTGA GAGTGCCGCT ACTCATCAAT AAATTTTTGA GCGCGGGTCC TAACGCTACT AACCTTTATT ACCATTTGAA ACGGGATTAT TCGCTTTATT TGGGGTATAA CTACACTTTT TAA	720 780 813
5	(2) INFORMATION FOR SEQ ID NO:72:	
10	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1317 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	•
	(ii) MOLECULE TYPE: DNA (genomic)	
15	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
20	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
25	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 11317</pre>	
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:72:	
30	ATGGCTTACA AACCTAACAA AAAGAAGTTA AAAGAATTAA GAGAGCAACC GAATTTATTT AGCATCTTAG ATAAGGCGA TGTTGCAACA AACAATCCTG TTGAAGAGTC AGACAAGGCC AATAAAATAC AAGAGCCACT CCCTTATGTC GTGAAAACGC AAATCAATAA AGCAAGCATG ATTTCTAGAG ATCCTATTGA ATGGGCAAAG TATTTAAGCT TTGAAAAACG AGTCTATAAG GATAATAGTA AAGAAGATGT CAATTTCTTT GCCAATGGTG AGATAAAAGA AAGTTCTCGTT GTTTATGAAG CGAATAAAGA AGGGTTTGAAA AGGCGCATCA CTAAAAAGATA CGATTTGATT	60 120 180 240 300 360
35	GATAGAAATA TTGATAGAAA TAGAGAATTT TTTATAAAAG AAATTGAAAT TCTAACCCAC ACAAACAGCT TAAAAGAATT GAAAGAGCAA GGGTTAGAAA TCCAATTGAC CCACCATAAT GAAACGCATA AGAAAGCCTT AGAAAATGGC AATGAAATCG TTAAAGAATA CGACCATCTT AAAGATATTT ACCAAGAAGT AGAAAGAACA AAAGATGGTG GATTGGTAAG AGAAATAATC CCCAGTATTT CTAGCGCTGA GTATTTCAAG CTTTACAACA AACTGCCTTT TGAATCAATA AACAATGAAA ATACCAAACT GAATACTAAC GACAATGAAG AAGTTAAAAA ACTAGAATTT	420 480 540 600 660 720
40	GAATTAGCTA AAGAAGTGCA TATTTTAATC CTAGAGCAAC AATTGCTTTC AGCAACAAAT TATTATTCTT GGATAGATAA AGATGATAAT GCGAATTTTG CTTGGAAAAT GCATAGGCTT ATCAATGAAA ATAAACTCAA AGAAAACCAT CTCAGCGCCA ATAACGCTAA TAAGATTAAG CAATTTTTCT TTAATAATG TTCTATTTTA GGCTGGACTA AAGAAGAACA AAGCGCTATA	780 840 900 960
45	CAAGAAACA GAGATTATTC TTTAAGAAGC GCTCTTTTAA GTTTAGAAGA AATCGCTCAA GCAAAAATTG AATTGCAAAA ATACTATGAA AGCGTTTATG TTAATGGTGA TGGGAATAAA AGAGAAATCA AGCCTTTTAA AGAAATTTTA AGAGACACCA ACAATTTTGA AAAAGCTTAT AAGGAGCGTT ATGACAAATT GGTAAGCTTG AGTGCAGCAA TCATTCAAGC TAAAGAGGGT GGTAATGAGC GACCAAATTC TAGTGCAAAT AACAATAACC CTATTAAAAA TACAATAGAG	1020 1080 1140 1200 1260
50	GGTAATGAGC GACCAAATTC TAGTGCAAAT AACAATAACC CTATTATATAAAAATAACAATAATAATAATAATAATAATA	1317
	(i) SEQUENCE CHARACTERISTICS:	

(A) LENGTH: 648 base pairs
(B) TYPE: nucleic acid

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	(C) STRANDEDNESS: double (D) TOPOLOGY: circular	
_	(ii) MOLECULE TYPE: DNA (genomic)	
. 5	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
10	(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
15	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1648</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:73:	-
20	ATGCAAGCGT TAAAATCATT GCTTGAAGTG ATTACAAAAC TCCAGAATCT AGGCGGCTAT TTGATGCATA TAGCTATTTT CATCATTTTT ATTTGGATTG GAGGGCTTAA GTTTGTGCCT TACGAAGCTG AAGGGATCGC CCCTTTTGTG GCCAACTCCC CTTTCTTTC TTTCATGTAT	60 120 180
	AAATTTGAAA AACCTGCATA CAAACAACAC AAAATGTCTG AATCCCAATC CATGCAAGAA GAAATGCAAG ATAACCCTAA AATCGTTGAA AACAAAGAAT GGCATAAAGA AAACCGCACT	240 300
25	TATTTAGTGG CTGAAGGTTT AGGGATTACG ATCATGATCC TAGGCATTTT GGTGCTTTTG	360
23	GGGCTTTGGA TGCCTTTAAT GGGCGTAGTT GGGGGCTTGC TTGTCGCTGG AATGACGATC ACCACCCTAT CTTTTTTATT CACAACGCCA GAAGTGTTTG TCAATCAGCA TTTCCCATGG	420 480
	CTTTCTGGGG CTGGAAGGCT AGTGGTTAAA GACTTGGCGT TATTTGCTGG AGGCTTGTTT	540
	GTGGCCGGAT TTGATGCGAA ACGCTATTTG GAGGGTAAAG GGTTTTGCTT GATGGACCGC	600
20	TCATCGGTAG GGATTAAAAC TAAATGCTCT AGCGGGTGTT GCTCTTAA	648
30	(2) INFORMATION FOR SEQ ID NO:74:	
	(_,,,,,,	
	(i) SEQUENCE CHARACTERISTICS:	
35	(A) LENGTH: 186 amino acids (B) TYPE: amino acid	
33	(D) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: protein	
40	(iii) HYPOTHETICAL: YES	
	(vi) ORIGINAL SOURCE:	
	(A) ORGANISM: Helicobacter pylori	
45	(ix) FEATURE:	
73	(A) NAME/KEY: misc feature	
	(B) LOCATION 1186	
	(wi) SPONENCE DESCRIPTION, SEC. ID NO. 24	
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:74:	
	Met Ile Lys Arg Ile Ala Cys Ile Leu Ser Leu Ser Ala Ser Leu Ala	
	1 5 10 15	
	Leu Ala Gly Glu Val Asn Gly Phe Phe Met Gly Ala Gly Tyr Gln Gln 20 25 30	
55	Gly Arg Tyr Gly Pro Tyr Asn Ser Asn Tyr Ser Asp Trp Arg His Gly	

						,°	3	40					45			
	Asn	Asp 50	35 Leu	Tyr	Gly				Lys	Leu	Gly	Phe 60		Gly	Phe	Ala
5	65	Lys				70			Tyr		75					80
,	Thr				85				Thr	90					95	
	-			100					Ile 105					110		
10	Gly	Leu	Ile 115	Gly	Gly	Val	Gln	Leu 120	Ala	Gly	Asn	Thr	Trp 125	Met	Phe	Pro
	туг	Asp	Val	Asn	Gln	Thr	Arg 135	Phe	Gln	Phe	Leu	Trp 140	Asn	Leu	Gly	Gly
15	Arg	Met	Arg	Val	Gly	Asp 150	Arg	Ser	Ala	Phe	Glu 155	Ala	Gly	Val	Lys	Phe 160
13	Pro	Met	Val	Asn	Gln 165	Gly	Ser	Lys	Asp	Val 170	Gly	Leu	Ile	Arg	Tyr 175	Tyr
	Ser	Trp	Tyr	Val 180	Asp	Tyr	Val	Phe	Thr 185	Phe						
20	(2)	INFO	ORMAT		FOR	SEQ	ID	NO : 75	5:							
		(i)	SEC	QUENC	CE CI	iara(CTER:	ISTIC	CS:							
25			(1	3) T	YPE:	ami	16 ar no ac line	cid	acio	is						
		(ii)) MOI	LECU	LE T	YPE:	pro	tein								
30		(iii)	HY:	POTH	ETIC	AL:	YES									
		(vi) OR:					icob	acte:	r py	lori					
35		(ix		A) N	AME/		mis		atur	e						
		(xi) SE	QUEN	CE D	ESCR	IPTI	ON:	SEQ	ID N	0:75	:				
40	Leu	Met	Arg	Ile	Ile	Ile	Arg	Leu	Leu		Phe	Lys	Met	Asn	Ala 15	Phe
	1 Leu	Lys	Leu		5 Leu	Ala	Ser	Leu		10 Gly	Gly	Leu	Trp	Tyr	-	Phe
45	Asn	Gly		20 Gly	Ser	Glu	Ile		25 Ala	Ile	Gly	Ile	Phe	30 Val	Leu	Ile
	Leu		35 Val	Phe	Phe	Ile		40 Pro	Val	Ser	Phe	Gln 60		Pro	Glu	Lys
	Arg	50 Glu	Glu	Tyr	Ile		55 Arg	Leu	Lys	Lys			Glu	Arg	Lys	Met 80
50	65 Ile	Leu	Gln	Asp		70 Gln	Lys	·Glu	Glu		75 Met	Arg	Leu	Tyr	Gln 95	Ala
	Lys	Lys	Glu			Ser	Arg	Gln			Asp	Leu	Lys	Glu 110	Gln	Met
55	Lys	Lys	Tyr	100 Ser					105	•						

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115 .

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	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO : 7	6 :							
5		(i	(A) L B) T	ENGT: YPE :	HARA H: 3 ami: OGY:	45 a no a	mino cid		ds						
10		(ii) MO	LECU	LE T	YPE:	pro	tein								
		(iii) HY	POTH	ETIC	AL:	YES									
15		(vi	-			OURC		icob	acte	r py	lori					
20	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1345 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:76:</pre>															
20		(xi) SE	QUEN	CE D	ESCR:	IPTI	ON:	SEQ	ID N	0:76	:				
	Met 1	Val	Lys	His	Tyr 5	Leu	Phe	Met	Ala	Val 10	Ser	Gln	Val	Phe	Phe 15	Ser
25	Phe	Phe	Leu	Val 20	Leu	Phe	Phe	Ile	Ser 25	Ser	Ile	Val	Leu	Leu 30	Ile	Ser
	Ile	Ala	Ser 35	Val	Thr	Leu	Val	Ile 40	Lys	Val	Ser	Phe	Leu 45	Asp	Leu	Val
30	Gln	Leu 50	Phe	Leu	Tyr	Ser	Leu 55	Pro	Gly	Thr	Ile	Phe 60	Phe	Ile	Leu	Pro
	Ile 65	Thr	Phe	Phe	Ala	Ala 70	Cys	Ala	Leu	Gly	Leu 75	Ser	Arg	Leu	Ser	Tyr 80
	Asp	His	Glu	Leu	Leu 85	Val	Phe	Phe	Ser	Leu 90	Gly	Val	Ser	Pro	Lys 95	Lys
35	Met	Thr	Lys	Ala 100	Phe	Val	Pro	Leu	Ser 105	Leu	Leu	Val	Ser	Ala 110	Ile	Leu
	Leu	Ala	Phe 115	Ser	Leu	Ile	Leu	Ile 120	Pro	Thr	Ser	Lys	Ser 125	Ala	Tyr	Tyr
40	Gly	Phe 130	Leu	Arg	Gln	Lys	Lys 135	Asp	Lys	Ile	Asp	Ile 140	Asn	Ile	Arg	Ala
	Gly 145	Glu	Phe	Gly	Gln	Lys 150	Leu	Gly	Asp	Trp	Leu 155	Val	Tyr	Val	Asp	Lys 160
	Thr	Glu	Asn	Asn	Ser 165	Tyr	Asp	Asn	Leu	Val 170	Leu	Phe	Ser	Asn	Lys 175	Ser
45	Leu	Ser	Gln	Glu 180	Ser	Phe	Ile	Leu	Ala 185	Gln	Lys	Gly	Asn	Ile 190	Asn	Asn
	Gln	Asn	Gly 195	Val	Phe	Glu	Leu	Asn 200	Leu	Tyr	Asn	Gly	His 205	Ala	Tyr	Phe
50	Thr	Gln 210	Gly	Asp	Lys	Met	Arg 215	Lys	Val	Asp	Phe	Glu 220	Glu	Leu	His	Leu
	225					Ser 230					235					240
	Gly	Thr	Asp	Tyr	245	Gly				250					255	
55	7	T	7	01 -	T	7 ~~	A ~~~	Dha	C	α 1 –	77-	T1 -	T 011	1727	802	Lan

55 Asn Lys Asn Gln Lys Arg Phe Ser Gln Ala Ile Leu Val Ser Leu

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				260					265					270		
			275	Ala				280					285	•	Ala	
5		290	Phe				295					300			Ala	
	305					310					315				Phe -	320
					325					Ala 330	Phe	Ile	Ser	Tyr	Leu 335	Leu
0	Phe	Arg	Lys	Phe 340	Ile	Leu	Lys	Arg	Tyr 345							
	(2)	INFO	RMAI	rion	FOR	SEQ	ID N	10:77	7:							
15		(i)	(<i>I</i>	QUENC A) LE B) TY	NGTH	I: 27 amir	76 an	nino cid		ls		•				
20		(ii)	MOI	LECUI	E TY	PE:	prot	cein								
	((iii)	HYI	РОТНЕ	TIC	AL: 3	ŒS									
25		(vi)		IGINA A) OF				icoba	actei	r py:	lori					
		(ix)		ATURI A) N		KEY:	mis	c fea	ature	e						
30				B) LO												
50		(xi)) SE	QUEN	CE DI	ESCR:	IPTI	: ис	SEQ :	ID N	0:77	:				
	1				5					10					Ile 15	
35	Trp	Leu	Tyr	Leu 20	Ala	Val	Phe	Phe	Leu 25	Leu	Ser	Val	Ser	Asp 30	Ala	Lys
	Glu	Ile	Ala 35	Met	Gln	Arg	Phe	Asp 40	Lys	Gln	Asn	His	Lys 45	Ile	Phe	Glu
40		50	Ala				55					60			Ala	
. •	65	Asn				70					75				Asp	80
	"Vål	Arg	Tyr	Asp	Thr 85	Lys	Thr	Lys	Glu	Ala 90	Leu	Leu	Glu	Gly	Asn 95	Ile
45				100					105					110		
			115	Asn				120					125		Val	
50		130	Val	Ser			135					140			Gly	
- -	145	Gln	Lys			150					155				Ser	TO
	Asp	Asn			165					170					175	
55	Gln	Lys	Ser	His	Leu	Ser	Met	Trp	Asn	Pro	Lys	Ile	Tyr	· Val	Gly	Ası

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180 185 Ile Pro Val Leu Tyr Leu Pro Tyr Ile Phe Met Ser Thr Ser Asn Lys 200 205 Arg Thr Thr Gly Phe Leu Tyr Pro Glu Phe Gly Thr Ser Asn Leu Asp 5 220 215 Gly Phe Ile Tyr Leu Gln Pro Phe Tyr Leu Ala Pro Lys Asn Ser Trp 230 235 Asp Met Thr Phe Thr Pro Gln Ile Arg Tyr Lys Arg Gly Phe Gly Leu 250 245 Asn Phe Glu Ala Arg Tyr Ile Asn Ser Lys Thr Gln Val Phe Ile Gln 10 265 Cys Ala Leu Phe 275 15 (2) INFORMATION FOR SEQ ID NO:78: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 224 amino acids (B) TYPE: amino acid 20 (D) TOPOLOGY: linear (ii) MOLECULE TYPE: protein (iii) HYPOTHETICAL: YES 25 (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori (ix) FEATURE: 30 (A) NAME/KEY: misc feature (B) LOCATION 1...224 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:78: 35 Met Ile Arg Leu Lys Gly Leu Asn Lys Thr Leu Lys Thr Ser Leu Leu Ala Gly Val Leu Leu Gly Ala Thr Ala Pro Leu Met Ala Lys Pro Leu Leu Ser Asp Glu Asp Leu Leu Lys Arg Val Lys Leu His Asn Ile Lys 40 Glu Asp Thr Leu Thr Ser Cys Asn Ala Lys Val Asp Gly Ser Gln Tyr Leu Asn Ser Gly Trp Asn Leu Ser Lys Glu Phe Pro Gln Glu Tyr Arg 70 45 Glu Lys Ile Phe Glu Cys Val Glu Glu Glu Lys His Lys Gln Ala Leu 90 Asn Leu Ile Asn Lys Glu Asp Thr Lys Asp Lys Glu Glu Leu Ala Lys 100 105 Lys Ile Lys Glu Ile Lys Glu Lys Ala Lys Val Leu Arg Gln Lys Phe 50 120 Met Ala Phe Glu Met Lys Glu His Ser Lys Glu Phe Pro Asn Lys Lys 135 Gln Leu Gln Thr Met Leu Glu Asn Ala Phe Asp Asn Gly Ala Glu Ser 150 155 55 Phe Ile Asp Asp Trp His Glu Arg Phe Gly Gly Ile Ser Arg Glu Asn

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					165					170					175	
		-	Lys	180	Leu				185					190		
5			Leu 195					200					205			
-	Lys	Lys 210	Ala	Leu	Met	Ile	Leu 215	Asp	Asn	Pro	Tyr	Leu 220	Leu	Trp	Leu	Val
10	(2)	INFO	RMAT	NOI	FOR	SEQ	ID N	10:79):							
10		(i)	(E) LE	ENGTH	I: 42 amir	TERI 29 am 10 ac 1ine	ino id		ls						
15		(ii)	MOL	ECUI	E TY	PE:	prot	ein							•	
	(ii) MOLECULE TYPE: protein(iii) HYPOTHETICAL: YES(vi) ORIGINAL SOURCE:(A) ORGANISM: Helicobacter pylori															
20		(vi)						.coba	cter	py]	Lori					
		(ix)	FEA			ŒY:	misc	_fea	ature	=						
25			(E	3) LO	CAT	ON :	14	129								
			SEÇ													
30	1		Tyr		5					10					15	
	Phe	Leu	Ile		Cys	Met	Ile	Asn		His	Ala	Lys	Ser		Leu	Pne
				20		_			25			_		30	D	
			Leu 35	Pro				40	Gln				45	Glu		Cys
35	Ser	Leu 50	35 Glu	Pro Cys	Leu	Lys	Asp 55	40 Leu	Gln Met	Leu	Gln	Asn 60	45 Gln	Glu Ile	Phe	Cys Ser
35	Ser Phe 65	Leu 50 Val	35 Glu Ser	Pro Cys Gln	Leu Tyr	Lys Asp 70	Asp 55 Asp	40 Leu Asn	Gln Met Asn	Leu Gln	Gln Asp 75	Asn 60 Glu	45 Gln Ser	Glu Ile Leu	Phe Lys	Cys Ser Thr 80
35 40	Ser Phe 65 Tyr	Leu 50 Val Tyr	35 Glu Ser Lys	Pro Cys Gln Asp	Leu Tyr Ile 85	Lys Asp 70 Leu	Asp 55 Asp Asn	40 Leu Asn Lys	Gln Met Asn Leu	Leu Gln Asn 90	Gln Asp 75 Pro	Asn 60 Glu Val	45 Gln Ser Phe	Glu Ile Leu Ile	Phe Lys Ala 95	Cys Ser Thr 80 Ser
	Ser Phe 65 Tyr	Leu 50 Val Tyr	35 Glu Ser Lys Pro	Pro Cys Gln Asp Ala 100	Leu Tyr Ile 85 Lys	Lys Asp 70 Leu Glu	Asp 55 Asp Asn Ser	40 Leu Asn Lys Tyr	Gln Met Asn Leu Glu 105	Leu Gln Asn 90 Pro	Gln Asp 75 Pro Lys	Asn 60 Glu Val Ile	45 Gln Ser Phe Glu	Glu Ile Leu Ile Leu 110	Phe Lys Ala 95 Ala	Cys Ser Thr 80 Ser Ile
40	Ser Phe 65 Tyr Gln Leu	Leu 50 Val Tyr Thr	35 Glu Ser Lys Pro	Pro Cys Gln Asp Ala 100 Lys	Leu Tyr Ile 85 Lys	Lys Asp 70 Leu Glu Val	Asp 55 Asp Asn Ser Val	Asn Lys Tyr Gly 120	Gln Met Asn Leu Glu 105 Arg	Leu Gln Asn 90 Pro	Gln Asp 75 Pro Lys Ala	Asn 60 Glu Val Ile	45 Gln Ser Phe Glu Leu 125	Glu Ile Leu Ile Leu 110 Val	Phe Lys Ala 95 Ala Met	Cys Ser Thr 80 Ser Ile Asn
	Ser Phe 65 Tyr Gln Leu Thr	Leu 50 Val Tyr Thr Leu Leu 130	35 Glu Ser Lys Pro Pro 115 Leu	Pro Cys Gln Asp Ala 100 Lys	Leu Tyr Ile 85 Lys Lys	Lys Asp 70 Leu Glu Val Leu	Asp 55 Asp Asn Ser Val Asn 135	Aon Lys Tyr Gly 120 Thr	Gln Met Asn Leu Glu 105 Arg	Leu Gln Asn 90 Pro Tyr Asn	Gln Asp 75 Pro Lys Ala Asn	Asn 60 Glu Val Ile Ile Asp 140	45 Gln Ser Phe Glu Leu 125 Phe	Glu Ile Leu Ile Leu 110 Val	Phe Lys Ala 95 Ala Met	Cys Ser Thr 80 Ser Ile Asn Gln
40	Ser Phe 65 Tyr Gln Leu Thr Val 145	Leu 50 Val Tyr Thr Leu Leu 130 Phe	35 Glu Ser Lys Pro Pro 115 Leu	Pro Cys Gln Asp Ala 100 Lys Ala Ser	Leu Tyr Ile 85 Lys Lys Tyr	Lys Asp 70 Leu Glu Val Leu Glu 150	Asp 55 Asp Asn Ser Val Asn 135 Glu	Asn Lys Tyr Gly 120 Thr	Gln Met Asn Leu Glu 105 Arg Arg	Leu Gln Asn 90 Pro Tyr Asn Glu	Gln Asp 75 Pro Lys Ala Asn Lys 155	Asn 60 Glu Val Ile Ile Asp 140 Leu	45 Gln Ser Phe Glu Leu 125 Phe	Glu Ile Leu Ile Leu 110 Val Asn Glu	Phe Lys Ala 95 Ala Met Ile	Cys Ser Thr 80 Ser Ile Asn Gln Tyr 160
40	Ser Phe 65 Tyr Gln Leu Thr Val 145 Lys	Leu 50 Val Tyr Thr Leu 130 Phe	35 Glu Ser Lys Pro 115 Leu Asp	Pro Cys Gln Asp Ala 100 Lys Ala Ser Glu	Leu Tyr Ile 85 Lys Lys Tyr Asp Lys 165	Lys Asp 70 Leu Glu Val Leu Glu 150 Glu	Asp 55 Asp Asn Ser Val Asn 135 Glu	Aon Lys Tyr Gly 120 Thr Ser Phe	Gln Met Asn Leu Glu 105 Arg Arg Pro	Leu Gln Asn 90 Pro Tyr Asn Glu Phe 170	Gln Asp 75 Pro Lys Ala Asn Lys 155 Ile	Asn 60 Glu Val Ile Ile Asp 140 Leu	45 Gln Ser Phe Glu Leu 125 Phe Glu Ala	Glu Ile Leu Ile Leu 110 Val Asn Glu Leu	Phe Lys Ala 95 Ala Met Ile Thr Leu 175	Cys Ser Thr 80 Ser Ile Asn Gln Tyr 160 Thr
40	Phe 65 Tyr Gln Leu Thr Val 145 Lys	Leu 50 Val Tyr Thr Leu 130 Phe Glu Glu	Ser Lys Pro Pro 115 Leu Asp Ile	Pro Cys Gln Asp Ala 100 Lys Ala Ser Glu Val 180	Leu Tyr Ile 85 Lys Lys Tyr Asp Lys 165 Glu	Lys Asp 70 Leu Glu Val Leu Glu 150 Glu Asn	Asp 55 Asp Asn Ser Val Asn 135 Glu Lys	Asn Lys Tyr Gly 120 Thr Ser Phe Leu	Gln Met Asn Leu Glu 105 Arg Arg Pro Pro Gln 185	Leu Gln Asn 90 Pro Tyr Asn Glu Phe 170 Asn	Gln Asp 75 Pro Lys Ala Asn Lys 155 Ile	Asn 60 Glu Val Ile Ile Asp 140 Leu Ile	45 Gln Ser Phe Glu Leu 125 Phe Glu Ala Ile	Glu Ile Leu Ile Leu 110 Val Asn Glu Leu Asn 190	Phe Lys Ala 95 Ala Met Ile Thr Leu 175	Cys Ser Thr 80 Ser Ile Asn Gln Tyr 160 Thr
40	Ser Phe 65 Tyr Gln Leu Thr Val 145 Lys Lys	Leu 50 Val Tyr Thr Leu 130 Phe Glu Glu Tyr	35 Glu Ser Lys Pro 115 Leu Asp	Pro Cys Gln Asp Ala 100 Lys Ala Ser Glu Val 180 Pro	Leu Tyr Ile 85 Lys Lys Tyr Asp Lys 165 Glu	Lys Asp 70 Leu Glu Val Leu Glu 150 Glu Asn	Asp 55 Asp Asn Ser Val Asn i35 Glu Lys Leu Asn	40 Leu Asn Lys Tyr Gly 120 Thr Ser Phe Leu Lys 200	Gln Met Asn Leu Glu 105 Arg Pro Pro Gln 185 Thr	Leu Gln Asn 90 Pro Tyr Asn Glu Phe 170 Asn Gln	Gln Asp 75 Pro Lys Ala Asn Lys 155 Ile Thr	Asn 60 Glu Val Ile Ile Asp 140 Leu Ile Thr	45 Gln Ser Phe Glu Leu 125 Phe Glu Ala Ile Asn 205	Glu Ile Leu Ile Leu 110 Val Asn Glu Leu Asn 190 His	Phe Lys Ala 95 Ala Met Ile Thr Leu 175 Thr	Cys Ser Thr 80 Ser Ile Asn Gln Tyr 160 Thr Pro

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												•				
		210					215					220				
	Gln			Met	Leu	Ala		Phe	Ile	Ser	Pro	_	Ser	Pro	Val	Ile
	225		1			230					235					240
	Glu	Tyr	Asp	Asp	Asp	Gly	Leu	Ile	Gly	Glu	Arg	Leu	Arg	Gln	Ile	Thr
5					245					250					255	
	Glu	Ser	Leu	Asn	Val	Glu	Val	Lys	His	Gln	Glu	Asn	Ile		Tyr	Lys
				260					265					270		_
	Gln	Ala		Ser	Phe	Ser	Lys		Phe	Arg	Lys	His	_	Ala	Phe	Phe
10	_	_	275	m\	7	77 -	T	280	mb	D	m\	77 1	285	~	01	T
10	гÀг	290	Ser	THE	Leu	Ile	295	ASII	Int	PIO	Inr	300	ьys	ser	GIY	reu
	Tla		Ser	Gln	Tle	Gly		Leu	Glu	Tvr	Lvs		Leu	Lvs	Tle	T.eu
	305	200	501			310				-1-	315			_,_		320
		Thr	Gln	Ile	Asn	Phe	Asn	Pro	Ser	Leu		Leu	Leu	Thr	Gln	
15					325					330					335	
	Lys	Asp	Arg	Lys	Asn	Leu	Phe	Ile	Val	Asn	Ala	Leu	Gln	Asn	Ser	Asp
				340					345					350		
	Glu	Thr		Ile	Glu	Tyr	Ala		Leu	Leu	Glu	Ser		Leu	Arg	His
20	.	m	355	3	M	C	C	360	-1 -	~1	T	~ 1	365	Db -	T	3
20	Asp	370	vai	ASII	Tyr	Ser	375	ALA	тте	GLY	Leu	380	Met	Pne	Leu	ASII
	Thr		Asp	Pro	His	Phe		Lvs	Ser	Phe	Gln		Ser	Leu	Glu	Asp
	385					390	-,-	-1-	``		395					400
	Asn	Gln	Val	Arg	Tyr	His	Asn	Gln	Ile	Tyr	Gln	Ala	Leu	Gly	Tyr	Ser
25					405					410					415	
	Phe	Glu	Pro	Ile	Lys	Asn	Glu	Ser	Glu	Thr	Lys	Lys	Glu			
				420		•			425							
	(2)	TATE	````````	TTON	EOD	CEO	TD .	TO - 9 (٠.							
30	(2)	TME	JRIM.	ITOM	FOR	SEQ	י ענ	10.50	<i>.</i>							
50		(i)	SEC	OUEN	CE CI	HARAC	CTER	STI	CS:							
		, ,	(2	A) LI	ENGT	H: 45	55 ar	nino	acio	is						
			(1	3) T	PE:	amir	no ac	cid								
			(1) T	OPOLO	OGY:	line	ear								
35																
		(11)	MOI	PECOI	E T	PE:	prot	eın								
		(iii)	HVI	ואירום	ያምቸ <i>ር</i> ያ	AL: }	ÆS			,						
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40		(vi)	OR	GINA	AL SO	OURCE	Ξ:									
						ISM:		.coba	acte	py]	ori					
		(ix)	FE	ATURE	E :								•			
46						ŒY:			ture	2						
45			(E	3) LC	CAT	ON 1	Ļ4	155								
		/ d \	ana	~~~~~		10 0 0 1	DOT			- D 170						
		(X1)	SEÇ	TOENC	E DE	ESCRI	PLIT	יאי :	יבט ו	א ענ	: 80					
	Val	Leu	Lve	Phe	Gln	Lys	Leu	Pro	Leu	Len	Phe	val	Ser	Ile	Leu	Tvr
50	1		_,,,	- 110	5	-,5			u	10					15	-1-
	_	Gln	Ser	Pro	Leu	Leu	Ala	Phe	Asp		Lys	Phe	Ser	Gly	_	Ala
				20					25	-	•			30		
	Glu	Ser	Val	Ser	Lys	Val	Gly	Phe	Asn	His	Ser	Lys	Leu	Asn	Ser	Lys
			35					40					45			_
55	~ 3 · ·	77.	T 7 -	D1	Th	CT 1	70 7	ml	51 -		_,	- 1 -	CTI lan aus	T7 ~	T	T

55 Glu Gly Ile Phe Pro Thr Ala Thr Phe Val Thr Ala Thr Ile Lys Leu

							55					60				
	~1 ~	50 Val	Acn	Ser	Asn	Leu		Pro	Lvs	Asn	Ile		Lys	His	Ser	Leu
	65					70					75					80
	Lvs	Ile	Gly	Val	Gly	Gly	Ile	Leu	Gly	Ala	Leu	Ala	Tyr	Asp	Ser	Thr
5	_				85					90					95	
	Lys	Thr	Leu	Ile	Asp	Gln	Ala	Thr		Gln	Ile	Tyr	Gly	Ser	GLu	Leu
			_	100	~1			m	105	Dho	T 011	Glv	λen	110	Pro	Trn
	Phe	Tyr		IIe	GТĀ	Arg	Trp	120	GIY	Pne	neu	GIY	125	Ala	110	115
10	T 1.5	7 cm	115 Ser	T.em	Tle	Glu	Ser		Ala	His	Thr	Arg		Tyr	Val	Leu
10	гур	130	DCI	200			135					140		_		
	Tyr	Asn	Ser	Tyr	Leu	Phe	Tyr	Ser	Tyr	Gly	Asp	Lys	Phe	His	Leu	Lys
	145					150					155					160
	Leu	Gly	Arg	Tyr		Ser	Asn	Met	Asp		Met	ser	ser	Tyr	175	GIII
15		-1	~ 1	7	165	Tra rac	Larg	T1_	Aen	170	Lve	Tle	Δla	Leu		Trp
	GLY	Pne	GIU	180	ASP	TYL	цуз	116	185	DCI	_,_			190	-2-	
	Phe	Ser	Ser		Gly	Arg	Ala	Leu		Phe	Gly	Gln	Trp	Ile	Arg	Asp
			195					200					205			
20	Trp	Tyr	Ala	Pro	Ile	Val		Glu	Asp	Gly	Arg		Glu	Val	Tyr	Asp
		210				~1 ·-	215		Dh.a	Com	C 0 x	220	Wie	Va 1	Gln	Val
		Ile	His	Ala	Ala	230	Leu	TYL	Pne	Ser	235	пуъ	nrs	Val		240
	225 Met	Pro	Phe	Ala	Tyr		Ser	Pro	Lys	Ile		Gly	Ala	Pro	Gly	Val
25					245					250					255	
	Lys	Ile	His	Ile	Asp	Ser	Asn	Pro		Phe	Lys	Gly	Leu	Gly	Leu	Arg
	_			260	-1 -	D	77 ~ 7	т1 о	265	Dro	ו בינו	ጥህም	Δla	270 Lys	Asp	Leu
	Ala	GIn	7nr 275	Thr	IIe	ASI	vai	280	Pile	PIO	var	- y -	285	2,2		
30	TVY	Asp	Val	Tvr	Trp	Arg	Asn		Lys	Ile	Gly	Glu	Trp	Gly	Ala	Ser
50		290					295					300				
	Leu	Leu	Ile	His	Gln	Arg	Phe	Asp	Tyr	Asn	Glu	Phe	Asn	Phe	Gly	Phe
	305				_	310		.	-1-	3	315	7	т1 о	G1 v	TTT	320 Tvr
25	Gly	Tyr	Tyr	GIn	Asn 325	Pne	GIY	Asn	Ата	330	ALA	Arg	116	Gly	335	- / -
35	Glv	Δen	Pro	Tle	Pro	Phe	Asn	Tvr	Arq		Asn	Ser	Val	Tyr	Gly	Gly
				340					345					350		
	Val	Phe	Ser	Asn	Ala	Ile	Thr	Ala	Asp	Ala	Val	Ser	Gly	Tyr	Val	Phe
			355					360					365			
40	Gly			Val	Tyr	Arg			Leu	Trp	GIY	380	Leu	Gly	Arg	ıyı
	ml	370	71-	The	7 ~~	7) a	375		Δrα	Ser	Tle			Asn	Leu	Gly
	385		MIA	1111	nr 9	390					395					400
	Tvr	Lys	Trp	Gly	Ser			Arg	Val	Asp	Val	Asn	Leu	Glu	Tyr	Tyr
45					405					410					415	
	Val	Val	Ser			Asn	Gly	Tyr			Asp	Tyr	Leu	Thr	GIY	Pro
		_		420		T	77 -	N ===	425		7 ~~	A ~~	Çar	430 Asn	Leu	Met
	Phe	Asn	Lys 435		rne	гÀг	ATG	440		GIU.	Asp	arg	445			Met
50	٧al	Ser			Phe	Phe	Phe									
20	, 44	450		-1-			455									

- (2) INFORMATION FOR SEQ ID NO:81:
- 55 (i) SEQUENCE CHARACTERISTICS:

- 151 -

```
(A) LENGTH: 282 amino acids
               (B) TYPE: amino acid
               (D) TOPOLOGY: linear
 5
         (ii) MOLECULE TYPE: protein
        (iii) HYPOTHETICAL: YES
         (vi) ORIGINAL SOURCE:
10
               (A) ORGANISM: Helicobacter pylori
         (ix) FEATURE:
               (A) NAME/KEY: misc feature
               (B) LOCATION 1...282
         (XI) SEQUENCE DESCRIPTION: SEQ ID NO:81:
     Met Gly Cys Ser Phe Ile Phe Lys Lys Val Arg Val Tyr Ser Lys Met
20
     Leu Val Ala Leu Gly Leu Ser Ser Val Leu Ile Gly Cys Ala Met Asn
                                     25
     Pro Ser Ala Glu Thr Lys Lys Pro Asn Asp Ala Lys Asn Gln Gln Pro
                                 40
     Val Gln Thr His Glu Arg Met Thr Thr Ser Ser Glu His Val Thr Pro
25
     Leu Asp Phe Asn Tyr Pro Val His Ile Val Gln Ala Pro Gln Asn His
     His Val Val Gly Ile Leu Met Pro Arg Ile Gln Val Ser Asp Asn Leu
                                         90
30
     Lys Pro Tyr Ile Asp Lys Phe Gln Asp Ala Leu Ile Asn Gln Ile Gln
                                     105
     Thr Ile Phe Glu Lys Arg Gly Tyr Gln Val Leu Arg Phe Gln Asp Glu
                                 120
     Lys Ala Leu Asn Val Gln Asp Lys Lys Ile Phe Ser Val Leu Asp
35
                             135
     Leu Lys Gly Trp Val Gly Ile Leu Glu Asp Leu Lys Met Asn Leu Lys
                         150
                                             155
     Asp Pro Asn Ser Pro Asn Leu Asp Thr Leu Val Asp Gln Ser Ser Gly
                                         170
40
     Ser Val Trp Phe Asn Phe Tyr Glu Pro Glu Ser Asn Arg Val Val His
                                     185
     Asp Phe Ala Val Glu Val Gly Thr Phe Gln Ala Ile Thr Tyr Thr Tyr
                                 200
     Thr Ser Thr Asn Asn Ala Ser Gly Gly Phe Asn Ser Ser Lys Ser Val
45
                             215
                                                 220
     Ile His Glu Asn Leu Asp Lys Asn Arg Glu Asp Ala Ile His Lys Ile
                        230
                                             235
     Leu Asn Arg Met Tyr Ala Val Val Met Lys Lys Ala Val Thr Glu Leu
                                         250
                     245
50
     Thr Lys Glu Asn Ile Ala Lys Tyr Arg Asp Ala Ile Asp Arg Met Lys
                                     265
                                                         270
    Gly Phe Lys Ser Ser Met Pro Gln Lys Lys
```

55 (2) INFORMATION FOR SEQ ID NO:82:

	(i)	(A	L) LE	NGTH PE:	: 28 amin	o am	ino id		.s						
	(ii)	MOI	ECUL	E TY	PE:	prot	ein								
(
	(vi)						.coba	ıcter	. bAJ	ori					
		(<i>I</i>	A) NA B) LC	ME/F	ON 1	L2	80								
	(xi)	SEÇ	QUENC	E DE	ESCRI	PTIC)N: 5	SEQ I	D NC	:82:					
1				5					10					15	
Ile			20					25					30		
		35					40					45			
	50					55					60				
65					70					75					80
				85					90					95	
_			100					105					110		
		115					120					125			
	130					135					140				
145					150					155					100
				165					170					1/5	
			180					185					190		
*		195					200					205			
	210					215					220				
225					230					235					240
Ile				245					250					255	
Arg	Glu	Lys		Gln	Glu	Glu	Arg	Glu 265	Asn	Glu	Tyr	Leu	Arg 270	Asn	Gl
Ile	Arg			Leu	Ser	Gly									
	Met 1 Ile Asn Glu Val 65 Val Gly Lys 145 Leu Asn Glu Glu Ser 225 Ile Arg	(iii) (iii) (vi) (ixi) (xi) Met Lys 1 Ile Leu Asn His Glu Ile 50 Val Lys 65 Val Leu Gly Asp Gly Ala Lys Gly 130 Lys Glu 145 Leu Gly Asn Asn Glu Lys Glu Tyr 210 Ser Asn 225 Ile Phe Arg Glu	(ii) MOI (iii) HYF (vi) ORI (ix) FEX (ix) SEQ (ix) SEQ (ix) SEQ Met Lys Leu 1	(A) LE (B) TY (D) TO (ii) MOLECUL (iii) HYPOTHE (vi) ORIGINA (A) OR (ix) FEATURE (A) NA (B) LO (xi) SEQUENC Met Lys Leu Arg 1 Ile Leu Ser Ala 20 Asn His Val Tyr 35 Glu Ile Pro Leu 50 Val Lys Leu Pro 65 Val Leu Thr Phe Gly Asp Ala Asn 100 Gly Ala Arg Ser 115 Lys Gly Val Val 130 Lys Glu Lys Pro 145 Leu Gly Ala Asn Asn Asn Lys Gln 180 Glu Lys Ile Asn 195 Glu Tyr Thr Thr 210 Ser Asn Arg Ala 225 Ile Phe Lys Asn Arg Glu Lys Leu 260	(A) LENGTH (B) TYPE: (D) TOPOLO (ii) MOLECULE TY (iii) HYPOTHETICA (vi) ORIGINAL SO (A) ORGANI (ix) FEATURE: (A) NAME/R (B) LOCATI (xi) SEQUENCE DE Met Lys Leu Arg Ala 1 1 5 1le Leu Ser Ala Cys 20 Asn His Val Tyr Thr 35 Glu Ile Pro Leu Asn 50 Val Lys Leu Pro Asn 65 Val Leu Thr Phe Lys 85 Gly Asp Ala Asn Lys 100 Gly Ala Arg Ser Asp 115 Lys Gly Val Val Met 130 Lys Glu Lys Pro Gln 145 Leu Gly Ala Asn Thr 165 Asn Asn Lys Gln Val 180 Glu Lys Ile Asn Gln 195 Glu Tyr Thr Thr Arg 210 Ser Asn Arg Ala Gly 225 Ile Phe Lys Asn Gly Arg Glu Lys Leu Gln 260 Ile Arg Ser Leu Leu	(A) LENGTH: 28 (B) TYPE: amin (D) TOPOLOGY: (ii) MOLECULE TYPE: (iii) HYPOTHETICAL: Y (vi) ORIGINAL SOURCE (A) ORGANISM: (ix) FEATURE: (A) NAME/KEY: (B) LOCATION I (xi) SEQUENCE DESCRI Met Lys Leu Arg Ala Ser 1 5 Ile Leu Ser Ala Cys Ser 20 Asn His Val Tyr Thr Pro 35 Glu Ile Pro Leu Asn Asp 50 Val Lys Leu Pro Asn Tyr 65 70 Val Leu Thr Phe Lys Leu 85 Gly Asp Ala Asn Lys Ile 100 Gly Ala Arg Ser Asp Ile 115 Lys Gly Val Val Met Ile 130 Lys Glu Lys Pro Gln Thr 145 Leu Gly Ala Asn Thr Lys 165 Asn Asn Lys Gln Val Ile 180 Glu Lys Ile Asn Gln Tyr 195 Glu Tyr Thr Thr Arg Lys 210 Ser Asn Arg Ala Gly Tyr 225 230 Ile Phe Lys Asn Gly Asn 245 Arg Glu Lys Leu Gln Glu 260 Ile Arg Ser Leu Leu Ser	(A) LENGTH: 280 am (B) TYPE: amino ac (D) TOPOLOGY: line (ii) MOLECULE TYPE: prot (iii) HYPOTHETICAL: YES (vi) ORIGINAL SOURCE: (A) ORGANISM: Heli (ix) FEATURE: (A) NAME/KEY: misc (B) LOCATION 12 (xi) SEQUENCE DESCRIPTION Met Lys Leu Arg Ala Ser Val 1	(A) LENGTH: 280 amino (B) TYPE: amino acid (D) TOPOLOGY: linear (ii) MOLECULE TYPE: protein (iii) HYPOTHETICAL: YES (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicoba (ix) FEATURE: (A) NAME/KEY: misc_fea (B) LOCATION 1280 (xi) SEQUENCE DESCRIPTION: SOURCE: (A) ORGANISM: Helicoba (xi) SEQUENCE DESCRIPTION: SOURCE: (B) LOCATION 1280 (xi) SEQUENCE DESCRIPTION: SOURCE: (A) NAME/KEY: misc_fea (B) LOCATION 1280 (xi) SEQUENCE DESCRIPTION: SOURCE: (A) ORGANISM: Helicoba (xi) SEQUENCE DESCRIPTION: SOURCE: (A) ORGANISM: Helicoba (B) LOCATION 1280 (xi) SEQUENCE DESCRIPTION: SOURCE: (A) ORGANISM: Helicoba (B) LoCATION 1280 (xi) SEQUENCE DESCRIPTION: SOURCE: (A) ORGANISM: Helicoba (B) LoCATION 1280 (xi) SEQUENCE DESCRIPTION: SOURCE: (A) ORGANISM: Helicoba (B) LoCATION 1280 (xi) SEQUENCE DESCRIPTION: Source (B) LoCATION 1280 (xi) SEQUENCE DESCRIPTION: Source (B) LoCATION 1280 (xi) SEQUENCE DESCRIPTION: Source (B) LoCATION 1280 (xi) ORIGINAL SOURCE: (A) ORGANISM: Helicoba (B) LoCATION 1280 (xi) ORIGINAL SOURCE: (A) ORGANISM: Helicoba (B) LoCATION 1280 (xi) ORIGINAL SOURCE: (A) ORGANISM: Helicoba (B) LoCATION 1280 (xi) ORIGINAL SOURCE: (A) ORGANISM: Helicoba (B) LoCATION 1280 (xi) ORIGINAL SOURCE: (A) ORGANISM: Helicoba (A) ORGANISM: Helicoba (B) LoCATION 1280 (xi) ORIGINAL SOURCE: (A) ORGANISM: Helicoba (B) LoCATION 1280 (xi) ORGANISM: Helicoba (A) ORGANISM: Helicoba (B) LOCATION 1280 (Xi) ORGANISM: Helicoba (A) ORGANICM: Helicoba (A) ORGANICM: Helicoba (A) ORGANICM: Helicoba (A) ORGANICM: Helicoba (A) ORGAN	(B) TYPE: amino acid (D) TOPOLOGY: linear (ii) MOLECULE TYPE: protein (iii) HYPOTHETICAL: YES (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter (ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1280 (xi) SEQUENCE DESCRIPTION: SEQ IN Met Lys Leu Arg Ala Ser Val Leu Ile 1	(A) LENGTH: 280 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (iii) MOLECULE TYPE: protein (iii) HYPOTHETICAL: YES (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pyl (ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1280 (xi) SEQUENCE DESCRIPTION: SEQ ID NO Met Lys Leu Arg Ala Ser Val Leu Ile Gly 1	(A) LENGTH: 280 amino acids (B) TYPE: amino acids (C) TOPOLOGY: linear (ii) MOLECULE TYPE: protein (iii) HYPOTHETICAL: YES (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori (ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1280 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:82: Met Lys Leu Arg Ala Ser Val Leu Ile Gly Val 1	(A) LENGTH: 280 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (iii) MOLECULE TYPE: protein (iii) HYPOTHETICAL: YES (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori (ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1280 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:82: Met Lys Leu Arg Ala Ser Val Leu Ile Gly Val Ala 1 5 10 Ile Leu Ser Ala Cys Ser Asn Tyr Ala Lys Lys Val 25 Asn His Val Tyr Thr Pro Val Tyr Asn Glu Leu Ile 35 60 Ual Lys Leu Pro Asn Tyr Lys Asp Thr Pro 50 Val Leu Thr Phe Lys Leu Val His His Ser Lys Lys 85 90 Gly Asp Ala Asn Lys Ile Leu Gln Tyr Lys Asn Tyr 100 Gly Ala Arg Ser Asp Ile Asp Phe Tyr Leu Gln Pro 115 120 Lys Gly Val Val Met Ile Ala Ser Asn Tyr Asn Asp 130 Lys Glu Lys Pro Gln Thr Phe Asp Val Leu Gln Gly 145 Leu Gly Ala Asn Thr Lys Asn Leu His Gly Tyr Asp 165 Asn Asn Lys Gln Val Ile Asn Glu Val Ala Arg Glu 195 Glu Lys Ile Asn Gln Tyr Tyr Lys Thr Leu Leu Gln 195 Glu Lys Ile Asn Gln Tyr Tyr Lys Thr Leu Leu Gln 195 Glu Lys Ile Asn Gln Tyr Tyr Lys Thr Leu Leu Gln 195 Glu Tyr Thr Thr Arg Lys Asn Asn Gln Arg Glu Ile 210 Ser Asn Arg Ala Gly Tyr Gln Met Arg Gln Asn Val 225 Ile Phe Lys Asn Gly Asn Leu Asn Met Gln Ala Lys 245 Arg Glu Lys Leu Gln Glu Glu Arg Glu Lys Leu Leu Gln Glu Tyr Gln Asn Glu Asn Glu Tyr 260 Ile Arg Ser Leu Leu Ser Gly Lys	(A) LENGTH: 280 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (ii) MOLECULE TYPE: protein (iii) HYPOTHETICAL: YES (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori (ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1280 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:82: Met Lys Leu Arg Ala Ser Val Leu Ile Gly Val Ala Ile 10 10 10 10 10 10 10 10 10 10 10 10 10	(A) LENGTH: 280 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (ii) MOLECULE TYPE: protein (iii) HYPOTHETICAL: YES (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori (ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1280 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:82: Met Lys Leu Arg Ala Ser Val Leu Ile Gly Val Ala Ile Leu 1	(A) LENGTH: 280 amino acids (B) TYPE: amino acid (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: protein (iii) HYPOTHETICAL: YES (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori (ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1280 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:82: Met Lys Leu Arg Ala Ser Val Leu Ile Gly Val Ala Ile Leu Cys 1

- 153 -

	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO : 8	3:							
5		(i	(A) L B) T	ENGT YPE :	H: 3 ami	CTER 93 a no a lin	mino cid		ds						
10) MO				pro YES	tein								
		(111	,													
15		(vi) OR (E: Hel	icob	acte	r py	lori					
		(ix		A) N	AME/		mis 1		atur	e						
20		(xi) SE	QUEN	CE D	ESCR	IPTI	ON:	SEQ	ID N	0:83	:				
	Met 1	Arg	Lys	Leu	Phe 5	Ile	Pro	Leu	Leu	Leu 10	Phe	Ser	Ala	Leu	Glu 15	Ala
25	Asn	Glu	Lys	Asn 20	Gly	Phe	Phe	Ile	Glu 25	Ala	Gly	Phe	Glu	Thr 30	Gly	Leu
	Leu	Glu	Gly 35	Thr	Gln	Thr	Gln	Glu 40	Lys	Arg	His	Thr	Thr 45	Thr	Lys	Asn
	Thr	Tyr 50	Ala	Thr	Tyr	Asn	Tyr 55	Leu	Pro	Thr	Asp	Thr 60	Ile	Leu	Lys	Arg
30	65					70	Asn				75					80
					85		Arg			90		_			95	
35				100			Pro	-	105					110		
			115				Asn	120		_		_	125			
	Ile	Pro 130	Lys	His	ser	rys	Ile 135	Val	Leu	Pro	GIY	G1u 140	Ala	Phe	Asp	ser
40	Leu 145	Lys	Ile	Asp	Pro	Tyr 150	Thr	Leu	Phe	Leu	Pro 155	Lys	Ile	Glu	Ala	Thr 160
	Ser	Thr	Ser	Ile	Ser 165	Asp	Ala	Asn	Thr	Gln 170	Arg	Val	Phe	Glu	Thr 175	Leu
45	Asn	Lys	Ile	Lys 180	Thr	Asn	Leu	Val	Val 185	Asn	Tyr	Arg	Asn	Glu 190	Asn	Lys
	Phe	Lys	Asp 195	His	Glu	Asn	His	Trp 200	Glu	Ala	Phe	Thr	Pro 205	Gln	Thr	Ala
	Glu	Glu 210	Phe	Thr	Asn	Leu	Met 215	Leu	Asn	Met	Ile	Ala 220	Val	Leu	Asp	Ser
50	Gln 225	Ser	Trp	Gly	Asp	Ala 230	Ile	Leu	Asn	Ala	Pro 235	Phe	Glu	Phe	Thr	Asn 240
		Pro	Thr	Asp	Cys 245		Asn	Asp	Pro	Ser 250		Cys	Val	Asn	Pro 255	Gly
	Thr	Asn	Gly	Leu	Val	Asn	Ser	Lys	Val	Asp	Gln	Lys	Tyr	Val	Leu	Asn

260

PCT/US97/19575

	Lys Glr	Asp 275	Ile	Val	Asn	Lys	Phe 280	Lys	Asn	Lys	Ala	Asp 285	Leu	Asp	Val
	Ile Val	Leu				295					300				
5	Pro Ser	Asn	Asn	Asp	Asp 310	Gly	Lys	His	Tyr	Gly 315	Gln	Leu	Gly	Val	Val 320
	Ala Sei	Ala		Asp 325	Pro	Lys	Lys	Leu	Phe 330	Gly	Asp	Asn	Leu	Lys 335	Thr
10	Ile Asr		Glu 340	Asp				345					350		
10	Lys Gly	355	Gly				360					365			
	Thr Lys	Asp	Gly	Gln	Val	Glu 375	Lys	Asp	Ser	Asn	Gly 380	Lys	Pro	Lys	Asp
15	Ser Ası 385	Gly	Leu	Pro	Tyr 390	Asn	Val	Cys							
	(2) IN	FORMAT	NOIT	FOR	SEQ	ID I	NO : 84	4:							
20	(:	i) SEQ	QUENC						is						
		(1	3) TY O) TO	PE:	ami	no a	cid								
25	(i	i) MOI	LECUI	LE T	YPE:	pro	tein								
	(ii	i) HY	POTHE	ETIC	AL:	YES									
30	(v	i) OR:	IGINA A) OF				icob	acte:	r py	lori					
	(i		ATURI A) NI B) LO	AME/				atur	e						
35	1×	i) SE						SEQ	ID N	O:84	:				
	Met Ly											Val	Leu	Ser	Ser
40	1 Ser Le			5					10					12	
40			20					25					30		
	Leu Gl	35					40					45			
45	Thr Gl					55					60				
	Asn Pr				70					75					00
	Ala Le			85					90					70	
50	Phe Ly		100	Asp				105	i				110	1	
	Gly Le	115	Asp	Tyr			120)				12:	•		
55	Pro As		Ile	Gln	Lev	135		: Val	. Sei	Tr	Gly 140	y Val	L Gly	Ser	: Asp

PCT/US97/19575

55

130

	Leu 145	Leu	Ala	Asp	Ile	Ile 150	Asp	Lys	Asp	Asn	Ala 155	Ser	Phe	Gly	Ile	Phe 160
	Gly	Gly	Val	Ala	Ile 165	Gly	Gly	Asn	Thr	Trp 170	Lys	Ser	Ser	Ala	Ala 175	Asn
5	Tyr	Trp	Lys	Glu 180	Gln	Ile	Ile	Glu	Ala 185	Lys	Gly	Pro	Asp	Val 190	Cys	Thr
	Pro	Thr	Tyr 195	Суз	Asn	Pro	Asn	Ala 200	Pro	Tyr	Ser	Thr	Asn 205	Thr	Ser	Thr
10	Val	Ala 210	Phe	Gln	Val	Trp	Leu 215	Asn	Phe	Gly	Val	Arg 220	Ala	Asn	Ile	Туг
	Lys 225	His	Asn	Gly	Val	Glu 230	Phe	Gly	Val	Arg	Val 235	Pro	Leu	Leu	Ile	Asn 240
		Phe	Leu	Ser	Ala 245	Gly	Pro	Asn	Ala	Thr 250	Asn	Leu	Tyr	Tyr	His 255	Leu
15	Lys	Arg	Asp	Tyr 260	Ser	Leu	Tyr	Leu	Gly 265	Tyr	Asn	Tyr	Thr	Phe 270	00	_
	(2)	INF	ORMA	rion	FOR	SEQ	ID I	NO : 8 !	5:							
20		(i)	(<i>I</i>	A) LI B) T	ENGTI YPE :	HARAG H: 14 amir DGY:	10 ar	mino cid		ls						
25		(ii)	MOI	LECUI	LE T	YPE:	prot	tein								
		(iii)	HYI	POTH	ETIC	AL: Y	YES									
30		(vi				OURCI		icoba	acte	r py	lori					
35		(ix)		A) N2	AME/I	KEY:		_	ature	2						
		(xi)	SEC	QUEN	CE DI	ESCRI	PTIC	ON: S	SEQ :	ID NO):85	:				
	Met 1	His	Pro	Ile	Met 5	Phe	Ala	Tyr	Ile	Ala 10	Asn	Ala	Leu	Ala	Gln 15	Ala
40		Lys	Ile	Asn 20		Thr	Leu	Cys	Met 25	Ala	Phe	Gln	Lys	Ile 30	Ser	Gln
	Val	Lys	Glu 35	Leu	Gly	Ile	Asp	Lys 40	Ala	Lys	Ser	Leu	Ile 45	Gly	Asn	Leu
45	Ser	Gln 50	Val	Ile	Ile	Tyr	Pro 55	Thr	Lys	Asp	Thr	Asp 60	Glu	Leu	Ile	Glu
	Cys 65	Gly	Val	Pro	Leu	Ser 70	Asp	Ser	Glu	Ile	Asn 75	Phe	Leu	His	Asn	Thr 80
	Asp	Met	Arg	Ala	Arg 85	Gln	Val	Leu	Val	Lys 90	Asn	Ile	Val	Thr	Asn 95	Ala
50	Ser	Ala	Phe	Ile 100	Glu	Ile	Asp	Leu	Lys 105	Lys	Ile	Cys	Lys	Asn 110	Tyr	Phe
	Ile	Phe	Leu 115	Ile	Ala	Met	Leu	Val 120	Ile	Glu	Lys	Ser	Ser 125	Met	Ile	Leu
	Lys	Lys	Gln	Thr	Lys	Lys	Leu	Ile	Arg	Lys	Ser	Ile				

(2) INFORMATION FOR SEQ ID NO:86:

5		(i)	(<i>I</i>	QUENC A) LE B) TY	NGTH PE:	[: 25 amir	66 an	ino cid		ls						
10		(ii)	MOI	LECUI	E TY	PE:	prot	ein								
10	((iii)	HYI	POTHE	TICA	T: }	ŒS									
15		(vi)		(GINA				icoba	acter	pyl	lori					
13	<pre>(ix) FEATURE:</pre>															
20		(xi)	SEC	QUENC	CE DE	ESCRI	PTIC	ON: S	SEQ]	D NO	0:86:	:				
	1				5					10					15	
25	Ala	Leu	Cys	Leu 20	Cys	Gly	Gly	Leu	Met 25	Ala	Glu	Gln	Asp	Pro 30	Lys	Glu
23	Leu	Ile	Phe	Ser	Gly	Ile	Thr	Ile 40		Thr	Asp	Lys	Asn 45	Phe	Thr	Arg
	Ala	Lys 50		Tyr	Phe	Glu	Lys 55	Ala	Cys	Lys	Ser	Asn 60	Asp	Ala	Asp	Gly
30	65	Ala		Leu		70					75					80
	Glu	Asn	Ala	Arg	Glu 85	Ser	Ile	Glu	Lys	Ala 90	Leu	Glu	His	Thr	Ala 95	Thr
35		_		Cys 100					105					110		
			115	Phe				120					125			
		130		Cys			135					140				
40		Phe	Tyr	Asn	Asp	Met 150	Ile	Lys	Gly	Leu	Lys 155	Lys	Asp	Lys	Lys	Asp 160
	145 Leu	Glu	Tyr	Tyr	Ser	Lys	Ala		Glu		Asn	Asn	Gly	Gly	Gly 175	
45	Ser	Lys	Leu	Gly 180	Gly	Asp	Tyr	Phe	Phe 185	Gly	Glu	Gly	Val	Thr 190	Lys	Asp
	Phe	Lys	Lys 195	Ala	Phe	Glu	Tyr	Ser 200	Ala	Lys	Ala	Cys	Glu 205	Leu	Asn	Asp
	Ala	Lys 210	Gly	Cys	Tyr	Ala	Leu 215	Ala	Ala	Phe	Tyr	Asn 220	Glu	Gly	Lys	Gly
50		Ala	Lys	Asp	Glu	Lys 230	Gln	Thr	Thr	Glu	Asn 235	Leu	Glu	Lys	Ser	Cys 240
	225 Lys	Leu	Gly	Leu	Lys 245		Ala	Cys	Asp	Ile 250		Lys	Glu	Gln	Lys 255	
55	(2)	INF	ORMA'	TION	FOR	SEQ	ID :	NO : 8	7:							

- 157 -

(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 242 amino acids (B) TYPE: amino acid 5 (D) TOPOLOGY: linear (ii) MOLECULE TYPE: protein (iii) HYPOTHETICAL: YES 10 (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori (ix) FEATURE: (A) NAME/KEY: misc feature 15 (B) LOCATION 1...242 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:87: Met Lys Lys Phe Phe Ser Gln Ser Leu Leu Ala Leu Ile Ile Ser Met 20 Asn Ala Val Ser Gly Met Asp Gly Asn Gly Val Phe Leu Gly Ala Gly 25 Tyr Leu Gln Gly Gln Ala Gln Met His Ala Asp Ile Asn Ser Gln Lys 25 Gln Ala Thr Asn Ala Thr Ile Lys Gly Phe Asp Ala Leu Leu Gly Tyr 55 Gln Phe Phe Phe Glu Lys His Phe Gly Leu Arg Leu Tyr Gly Phe Phe 70 30 Asp Tyr Ala His Ala Asn Ser Ile Lys Leu Lys Asn Pro Asn Tyr Asn 85 90 Ser Glu Ala Ala Gln Val Ala Ser Gln Ile Leu Gly Lys Gln Glu Ile 105 Asn Arg Leu Thr Asn Ile Ala Asp Pro Arg Thr Phe Glu Pro Asn Met 35 120 Leu Thr Tyr Gly Gly Ala Met Asp Val Met Val Asn Val Ile Asn Asn 135 140 Gly Ile Met Ser Leu Gly Ala Phe Gly Gly Ile Gln Leu Ala Gly Asn 155 150 40 Ser Trp Leu Met Ala Thr Pro Ser Phe Glu Gly Ile Leu Val Glu Gln 170 Ala Leu Val Ser Lys Lys Ala Thr Ser Phe Gln Phe Leu Phe Asn Val 185 Gly Ala Arg Leu Arg Ile Leu Lys His Ser Ser Ile Glu Ala Gly Val 45 195 200 Lys Phe Pro Met Leu Lys Lys Asn Pro Tyr Ile Thr Ala Lys Asn Leu 215 Asp Ile Gly Phe Arg Arg Val Tyr Ser Trp Tyr Val Asn Tyr Val Phe 225 230 235 50 Thr Phe

- (2) INFORMATION FOR SEQ ID NO:88:
- 55 (i) SEQUENCE CHARACTERISTICS:

			(B) TY	PE:	: 26 amin GY:	o ac	id	acid	s						
5		(ii)	MOL	ECUL	E TY	PE:	prot	ein								
	(iii)	HYF	OTHE	TICA	L: Y	ES									
10		(vi)	ORI			URCE		.coba	ıcter	pyl	ori.					
15		(ix)) NA	ME/K	EY:			ture	:						
13		(xi)	SEÇ	QUENC	E DE	SCRI	PTIC	ON: S	SEQ 3	D NC	:88:					
	Met 1	Asn	Tyr	Pro	Asn 5	Leu	Pro	Asn	Ser	Ala 10	Leu	Glu	Ile	Ser	Glu 15	Gln
20	Pro	Glu	Val	Lys 20	Glu	Ile	Thr	Asn	Glu 25	Leu	Leu	Lys	Gln	Leu 30	Gln	Asn
	Ala	Leu	Arg 35	Ser	Asn	Ala	His	Phe 40	Ser	Glu	Gln	Val	Glu 45	Leu	Ser	Leu
25	_	50	Ile				55					60			Phe	
	65	Asn				70					75				Glu	80
	Leu				85					90					95	Glu
30				100					105					110	Gln	
			115					120					125			Ile
35	_	130					135					140				Thr
	145					150					155					Gln 160
					165					170					Ala 175	
40				180					185					190		Thr
			195					200					205			Asn
45	Gln	Thr 210		Ala	Ile	Thr	Asn 215		Asn	Glu	Ala	Lys 220	Glu	Ser	Ala	Thr
15	Thr 225	Gln	Ile	Asn	Ala	Asn 230		Gln	Glu	Ala	Ile 235	Asn	Asn	Ile	Thr	Gln 240
	Glu	Lys	Thr	Gln	Ala 245	Thr		Glu	Ile	Thr 250	Glu	Ala	Lys	Lys	Thr 255	Asp
50	His	Tyr	Gln	Asn 260	Ile		Phe	Phe	Glu 265	Phe						

- (2) INFORMATION FOR SEQ ID NO:89:
- 55 (i) SEQUENCE CHARACTERISTICS:

- 159 -

```
(A) LENGTH: 544 amino acids
               (B) TYPE: amino acid
               (D) TOPOLOGY: linear
 5
         (ii) MOLECULE TYPE: protein
        (iii) HYPOTHETICAL: YES
         (vi) ORIGINAL SOURCE:
10
               (A) ORGANISM: Helicobacter pylori
         (ix) FEATURE:
               (A) NAME/KEY: misc_feature
               (B) LOCATION 1...544
15
        (xi) SEQUENCE DESCRIPTION: SEQ ID NO:89:
     Val Ile Glu Thr Ile Pro Lys His Ser Lys Ile Val Leu Pro Gly Glu
                                         10
20
     Ala Phe Asp Ser Leu Lys Glu Ala Phe Asp Lys Ile Asp Pro Tyr Thr
     Phe Phe Pro Lys Phe Glu Ala Thr Ser Thr Ser Ile Ser Asp Thr
                                 40
     Asn Thr Gln Arg Val Phe Glu Thr Leu Asn Asn Ile Lys Thr Asn Leu
25
     Ile Met Lys Tyr Ser Asn Glu Asn Pro Asn Asn Phe Asn Thr Cys Pro
                                             75
                        70
     Tyr Asn Asn Asn Gly Asn Thr Lys Asn Asp Cys Trp Gln Asn Phe Thr
     Pro Gln Thr Ala Glu Glu Phe Thr Asn Leu Met Leu Asn Met Ile Ala
30
                                     105
     Val Leu Asp Ser Gln Ser Trp Gly Asp Ala Ile Leu Asn Ala Pro Phe
                                 120
     Glu Phe Thr Asn Ser Ser Thr Asp Cys Asp Ser Asp Pro Ser Lys Cys
35
                             135
                                                 140
     Val Asn Pro Gly Val Asn Gly Arg Val Asp Thr Lys Val Asp Gln Gln
                         150
                                             155
     Tyr Ile Leu Asn Lys Gln Gly Ile Ile Asn Asn Phe Arg Lys Lys Ile
                                         170
40
     Glu Ile Asp Ala Val Val Leu Lys Asn Ser Gly Val Val Gly Leu Ala
                                     185
                                                         190
     Asn Gly Tyr Gly Asn Asp Gly Glu Tyr Gly Thr Leu Gly Val Glu Ala
                                 200
     Tyr Ala Leu Asp Pro Lys Lys Leu Phe Gly Asn Asp Leu Lys Thr Ile
45
                            215
                                                 220
     Asn Leu Glu Asp Leu Arg Thr Ile Leu His Glu Phe Ser His Thr Lys
                        230
                                            235
    Gly Tyr Gly His Asn Gly Asn Met Thr Tyr Gln Arg Val Pro Val Thr
                     245
                                        250
50
    Lys Asp Gly Gln Val Glu Lys Asp Ser Asn Gly Lys Pro Lys Asp Ser
                                     265
    Asp Gly Leu Pro Tyr Asn Val Cys Ser Leu Tyr Gly Gly Ser Asn Gln
                                 280
```

Pro Ala Phe Pro Ser Asn Tyr Pro Asn Ser Ile Tyr His Asn Cys Ala

300

295

	_	Val	Pro	Ala	Gly	Phe 310	Leu	Gly	Val	Thr	Ala 315	Ala	Val	Trp	Gln	Gln 320
	305 Leu	Ile	Asn	Gln	Asn 325		Leu	Pro	Ile	Asn 330	Tyr	Ala	Asn	Leu	Gly 335	Ser
5	Gln	Thr	Asn	Tyr 340		Leu	Asn	Ala	Ser 345		Asn	Thr	Gln	Asp 350	Leu	Ala
	Asn	Ser		Leu	Ser	Thr	Ile	Gln 360		Thr	Phe	Val	Thr 365	Ser	Ser	Val
10	Thr		355 His	His	Phe	Ser	Asn 375		Ser	Gln	Ser	Phe 380		Ser	Pro	Ile
10		370 Gly	Val	Asn	Ala	Lys 390		Gly	Tyr	Gln	Asn 395	-	Phe	Asn	Asp	Phe 400
	385 Ile	Gly	Leu	Ala	Tyr 405		Gly	Ile	Ile	Lys 410		Asn	Tyr	Ala	Lys 415	Ala
15	Val	Asn	Gln	Lys 420	Val	Gln	Gln	Leu	Ser 425		Gly	Gly	Gly	Ile 430	Asp	Leu
	Leu	Leu	Asp			Thr	Thr	Tyr 440		Asn	Lys	Asn	Ser	Pro	Thr	Gly
20	Ile	Gln 450	Thr	Lys	Arg	Asn	Phe 455		Ser	Ser	Phe	Gly 460	Ile	Phe	Gly	Gly
20	Leu 465	Arg	Gly	Leu	Tyr	Asn 470		Tyr	Tyr	Val	Leu 475			Val	Lys	Gly 480
	Ser	Gly	Asn	Leu	Asp	Val	Ala	Thr	Gly	Leu 490		туг	Arg	Tyr	Lys 495	His
25	Ser	Lys	Tyr	Ser 500			Ile	Ser	11e 505			Ile	Gln	Arg 510	Lys	Ala
	Ser	Val	Val 515	Ser	Ser	Gly	Gly	Asp 520	Tyr	Thr	Asn	Ser	Phe 525	Val	Phe	Asn
30	Glu	Gly 530	Ala	Ser	His	Phe	Lys 535			Phe	Asn	Tyr 540	Gly	Gly	Cys	Phe
	(2)	INF	ORMA'	TION	FOR	SEQ	ID :	NO : 9	0:							
35		(i	(A) L B) T	ENGT YPE :	H: 3 ami	CTER 56 a no a lin	mino cid		ds						
40		(ii) MO	LECU	LE T	YPE:	pro	tein								
40		(iii) HY	POTH	ETIC	AL:	YES									
15		(vi) OR					icob	acte	r py	lori					
45		(ix		A) N	AME/		mis	_	atur	e						
50		(xi) SE	QUEN	CE D	ESCR	IPTI	ON:	SEQ	ID N	10:90	·:				
	Leu 1	Met	Lys	Ser	Ile 5	Leu	Leu	Phe	Met	Ile 10	. Phe	. Val	. Val	Cys	Gln 15	Lev
55	Glu	Gly	Lys	Lys 20		Ser	Gln	Asp	Asn 25		Lys	. Val	Asp	Туr 30	· Asn	Туі

	Tyr	Leu	Arg 35	Lys	Gln	Asp	Leu	His 40	Ile	Ile	Lys	Thr	Gln 45	Asn	Asp	Leu
	Ser	Asn 50	Ala	Trp	Tyr	Leu	Pro 55	Pro	Gln	Lys	Ala	Pro 60	Lys	Glu	His	Ser
5	Trp 65	Val	Asp	Phe	Ala	Lys 70	Lys	Tyr	Leu	Asn	Met 75	Met	Asp	Tyr	Leu	Gly 80
	Thr	Tyr	Phe	Leu	Pro 85	Phe	Tyr	His	Ser	Phe 90	Thr	Pro	Ile	Phe	Gln 95	Trp
10	Tyr	His	Pro	Asn 100		Asn	Pro	Tyr	Gln 105	Arg	Asn	Glu	Phe	Lys 110	Phe	Gln
	Ile	Ser	Phe 115	Arg	Val	Pro	Val	Phe 120	Arg	His	Ile	Leu	Trp 125	Thr	Lys	Gly
	Thr	Leu 130	Tyr	Leu	Ala	Tyr	Thr 135	Gln	Thr	Asn	Trp	Phe 140	Gln	Ile	Tyr	Asn
15	Asp 145	Pro	Gln	Ser	Ala	<u>Pro</u>	Met	Arg	Met	Ile	Asn 155	Phe	Met	-Pro	Glu	Leu 160
	Ile	Tyr	Val	Tyr	Pro 165	Ile	Asn	Phe	Lys	Pro 170	Phe	Gly	Gly	Lys	Ile 175	Gly
20	Asn	Phe	Ser	Glu 180	Ile	Trp	Ile	Gly	Trp 185	Gln	His	Ile	Ser	Asn 190	Gly	Val
	Gly	Gly	Ala 195	Gln	Cys	Tyr	Gln	Pro 200	Phe	Asn	Lys	Glu	Gly 205	Asn	Pro	Glu
	Asn	Gln 210	Phe	Pro	Gly	Gln	Pro 215	Val	Ile	Val	Lys	Asp 220	Tyr	Asn	Gly	Gln
25	Lys 225	Asp	Val	Arg	Trp	Gly 230	Gly	Cys	Xaa	Ser	Val 235	Xaa	Xaa	Gly	Asn	Xaa 240
	Leu	Cys	Phe	Val	Leu 245	Val	Trp	Glu	Lys	Gly 250	Gly	Leu	Lys	Ile	Met 255	Val
30	Ala	Tyr	Trp	Pro 260	Tyr	Val	Pro	Tyr	Asp 265	Gln	Ser	Asn	Pro	Gln 270	Leu	Ile
	Asp	Tyr	Met 275	Gly	Tyr	Gly	Asn	Ala 280	Lys	Ile	Asp	Tyr	Arg 285	Arg	Gly	Arg
	His	His 290	Phe	Glu	Leu	Gln	Leu 295	Tyr	Asp	Ile	Phe	Thr 300	Gln	Tyr	Trp	Arg
35	Tyr 305	Asp	Arg	Trp	His	Gly 310	Ala	Phe	Arg	Leu	Gly 315	Tyr	Thr	Tyr	Arg	Ile 320
	Asn	Pro	Phe	Val	Gly 325	Ile	Tyr	Ala	Gln	Trp 330	Phe	Asn	Gly	Tyr	Gly 335	Asp
40	Gly	Leu	Tyr	Glu 340	Tyr	Asp	Val	Phe	Ser 345	Asn	Arg	Ile	Gly	Val 350	Gly	Ile
	Arg	Leu	Asn 355	Pro												

(2) INFORMATION FOR SEQ ID NO:91:

45

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 675 amino acids
 - (B) TYPE: amino acid
 - (D) TOPOLOGY: linear

- (ii) MOLECULE TYPE: protein
- (iii) HYPOTHETICAL: YES
- 55 (vi) ORIGINAL SOURCE:

PCT/US97/19575

(A) ORGANISM: Helicobacter pylori

(ix) FEATURE:

(A) NAME/KEY: misc_feature

5 (B) LOCATION 1...675

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:91:

		(xi)	SEC	OMENC	E DE	SCRI	PTIC)N: 5	EQ 1	טא ט.	: 91:					
10	1				5	Ser				10					15	
10	Leu	Ile	Val	Ile 20	Val	Cys	Val	Ser	Ile 25	Leu	Gly	Val	Ser	Leu 30	Asn	Ser
	Arg	Val	Lys 35	Glu	Ile	Leu	Lys	Glu 40	Ser	Ala	Leu	His	Ser 45	Met	Gln	Asp
15	Ser		His	Phe	Lys	Val	Lys 55		Val	Gln	Ser	Val 60	Leu	Glu	Asn	Thr
		50 Thr	Ser	Met	Gly	Ile 70	Val	Lys	Glu	Met	Leu 75	Pro	Glu	Asp	Thr	Lys 80
20	65 Arg	Glu	Ile	Lys	Ile 85	Gln	Leu	Leu	Lys	Asn 90	Phe	Ile	Leu	Ala	Asn 95	Ser
20	His	Val	Ala	Gly 100		Ser	Met	Phe	Phe 105	Lys	Asp	Arg	Glu	Asp 110	Leu	Arg
	Leu	Thr	Leu 115	Leu	Arg	Asp	Asn	Asp 120	Thr	Ile	Lys	Leu	Met 125	Glu	Asn	Pro
25	Ser	Leu 130	Gly	Ser	Asn	Pro	Leu 135	Ala	Gln	Lys	Ala	Met 140	Lys	Asn	Lys	Glu
	Ile	Ser	Lys	Ser	Leu	Pro	Tyr	Tyr	Arg	Lys	Met 155	Pro	Asn	Gly	Ala	Glu 160
	145		_	_		150	_	•	D	T		T	C1.,	λαπ	Thr	
	Val	Tyr	Gly	Val		Ile	Leu	Leu	Pro		Pne	гур	GIU	ASII	175	
30			_		165	_	30 - L	+ 1 -	Db -	170	C	T10	λαη	Ser		Ser
	Glu	Val	Val		Val	Leu	Mec	тте	185	Pne	Ser	116	YSP	190		
			-1.	180	*	Asn	7~~	gar		T.em	Phe	Leu	Ile	_	Val	Lys
	Asn	GIu	11e	Thr	гув	ASII	Arg	200	ASP	n-u	1110		205	1		•
35	C1.,	Tare	133 133	T.e.u	Leu	Ser	Ala		Lvs	Ser	Leu	Gln	Asp	Lys	Ser	Ile
55	_	210					215					220				
	Thr	Glu	Ile	Tyr	Lys	Ser	Val	Pro	Lys	Ala	Thr	Asn	Glu	Val	Met	Ala
	225					230					235					240
40					245	Ser				250					255	
				260					265					270		Gly
			275					280					285			Ile
45		290					295					300				Val
	Val	Ala	Ala	Ser	Ala	Ile	Met	Val	Leu	Ala	Leu	Ile	Ile	Ala	Ile	Thr
	305					310					315		3			320
50					325					330					335	
	Thr	Leu	Ser	His		Phe	Lys	Leu	Leu 345	Asn	Asn	Gln	Ala	His 350	Ser	Ser
	Asp	Ile	Lys	Leu	Val	Glu	Ala	Arg		Asn	Asp	Glu	Leu 365	Gly	Arg	Met
55	Gln	Thr	Ala	Ile	Asn	Lys	Asn			Gln	Thr	Gln	. Lys	Thr	Met	Gln

- 163 -

	93	370		<i>α</i> 1-	7.1	17-1	375		The sec	T1 -	T	380	**- 1	C	3	170 1
		_	Arg	GIII	ALG	Val 390	GIII	Asp	IIIL	TIE	395	vaı	vai	ser	Asp	400
	385		Gl v	Δen	Dhe	Ala	Va 1	Δνα	Tla	Thr		Glu	Bro	Λ 1 =	Sar	
5	пуз	ALA	GIY	ASII	405	AIG	Val	ALG	116	410	Ата	GIU	PLO	AIA	415	PIO
,	λcn	T.011	Lve	Glu		Arg	Agn	Δla	T.e.ii		Gly	Tla	Mat) en		T.Au
	Asp	Deu	шуз	420	Deu	Arg	ASP	AIG	425		Gry	116	Mec	430	TYL	Бец
	Gl n	Glu	Sar		Glv	Thr	His	Met			Tla	Dha	Lare		Dha	Glu
	GIII	GIU	435	VUL	Cly		*****	440	110	JCI	110	FIIC	445	116	FIIÇ	GIU
10	Sar	ጥኒታ		Glv	Leu	Asp	Phe		Glv	Ara	Tle	Gln		Δla	Ser	Glv
10	361	450	001	01,			455	•	- -1	9		460	1141	n.u	501	O ₁
	Ara		Glu	Leu	Val	Thr		Ala	Leu	Glv	Gln		Ile	Gln	Lvs	Met
	465					470				1	475				-1-	480
		Glu	Thr	Ser	Ser	Asn	Phe	Ala	Lvs	Asp		Ala	Asn	Asp	Ser	
15					485				•	490					495	
	Asn	-Leu	Lys	-Glu-	Cys	Val	Gln	Asn	Leu	Glu	Lys	Ala	Ser	Asn	Ser	Gln
			_	500	-				505		•			510		
	His	Lys	Ser	Leu	Met	Glu	Thr	Ser	Lys	Thr	Ile	Glu	Asn	Ile	Thr	Thr
			515					520					525			
20	Ser	Ile	Gln	Gly	Val	Ser	Ser	Gln	Ser	Glu	Ala	Met	Ile	Glu	Gln	Gly
		530					535					540				
	Lys	Asp	Ile	Lys	Ser	Ile	Val	Glu	Ile	Ile	Arg	Asp	Ile	Ala	Asp	Gln
	545					550					555					560
0.5	Thr	Asn	Leu	Leu		Leu	Asn	Ala	Ala		Glu	Ala	Ala	Arg		Gly
25				_	565					570					575	_
	Glu	His	Gly		Gly	Phe	Ala	Val		Ala	Asp	Glu	Val	_	Lys	Leu
	- 1 -	~1	•	580	a1	•	a	.	585	~1	-1.	~ 1		590	71 -	3
	Ala	GIU	_	THE	GIN	Lys	ser		ser	GIU	TTE	GIU		ASN	TTE	Asn
30	T1.0	T 011	595	C1 =	602	Ile	Ca*	600	mp	C	<i>c</i> 1	C ~ ~	605	Tara) CD	Gl n
50	176	610	Val	GIII	361	116	615	ASP	1111	261	GIU	620	TIE	Lys	VOII	GIII
	Va 1		Glu	Val	Glu	Glu		Asn	Ala	Ser	Tle		Ala	Leu	Arg	Ser
	625	-,-				630		.,			635		•••		5	640
		Thr	Glu	Gly	Asn	Leu	Lvs	Ile	Ala	Ser		Ser	Leu	Glu	Ile	Ser
35				•	645		•			650					655	
	Gln	Glu	Ile	Asp	Lys	Val	Ser	Asn	Asp	Ile	Leu	Glu	Asp	Val	Asn	Lys
				660	_				665				_	670		
	Lys	Gln	Phe													
			675													
40																
	(2)	INFO	RMAI	NOI	FOR	SEQ	ID N	10:92	:							
		(i)		•		IARAC										
4.5				-		[: 27		-	acid	ls						
45						amin										
			(I)) TC	POLC	GY:	line	ear								
		12.15														
		(11)	MOL	ECUL	E TY	PE:	prot	eın								
50		4223	*****	Omtre	·m - ~ -	.	rnc									
JU	0 (iii) HYPOTHETICAL: YES															
		(254.)	OD T	CTNIA	T. GO	URCE	• .									
		(4 T)				SM:		coho	ct a=		ori					
			(24	., OR	.GMIN I	: ויזכ	 1	.copa	ccer	ьλт	Ur1					

55 (ix) FEATURE:

ICDOCID: JMO 001022281 1 -

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(A) NAME/KEY: misc_feature
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(B) LOCATION 1...271

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:92:

		(xi)	SEC	QUENC	E DE	SCRI	PTIC	ON: 5	SEQ 3	ED NO):92:					
5																_
	1				5					10				Val	15	
	Phe			20					25					Lys 30		
10	His	Lys	Ile 35	Thr	Arg	Glu	Leu	Lys 40	Val	Gly	Ala	Asn	Pro 45	Val	Pro	His
	Ala	Gln 50	Ile	Leu	Gln	Ser	Val 55	Val	Asp	Asp	Leu	Lys 60	Glu	Lys	Gly	Ile
15	65	Leu				70					75			`Asn		80
15	Leu				85					90				Arg	95	
		_		100					105					Gly 110		
20			115					120					125	Ile		
		130					135					140		Asn		
	Ala	Asn	Gln	Gly	Arg	Ala	Leu	Ile	Leu	Leu	His	Lys	Gln	Gly	Leu	Ile
25	145					150					155			_	-1-	160
					165					170				Asp	175	
	-			180					185					Ala 190		
30			195					200					205			тут
		210					215					220		Asp		
	Ser	Pro	Tyr	Ala	Asn	Leu	Val	Ala	Ser	Arg	Glu	Asp	Asn	Ala	Gln	Asp
35	225					230					235					240
					245					250				Lys	255	
	Lys	Phe	Ile	Leu 260		Thr	Tyr	Lys	Gly 265	Ala	Ile	Ile	Pro	Ala 270	Phe	
40						CEO	TD	N() - 0	2.							
	(2)					SEQ										
		(i				HARA H: 1				ds						
45				B) T	YPE:	ami	no a	cid								

- (B) TYPE: amino acid
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein
- 50 (iii) HYPOTHETICAL: YES
 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Helicobacter pylori
- 55 (ix) FEATURE:

PCT/US97/19575

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(A) NAME/KEY: misc_feature

(B) LOCATION 1...161

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:93:

5 Met Phe Phe Lys Thr Tyr Gln Lys Leu Leu Gly Ala Ser Cys Leu Ala Leu Tyr Leu Val Gly Cys Gly Asn Gly Gly Gly Glu Ser Pro Val Glu Met Ile Ala Asn Ser Glu Gly Thr Phe Gln Ile Asp Ser Lys Ala 10 Asp Ser Ile Thr Ile Gln Gly Val Lys Leu Asn Arg Gly Asn Cys Ala 55 Val Asn Phe Val Pro Val Ser Glu Thr Phe Gln Met Gly Val Leu Ser 15 75 Gln Val Thr Pro Ile Ser Ile Gln Asp Phe Lys Asp Met Ala Ser Thr 90 Tyr Lys Ile Phe Asp Gln Lys Lys Gly Leu Ala Asn Ile Ala Asn Lys 105 20 Ile Ser Gln Leu Glu Gln Lys Gly Val Met Met Glu Pro Gln Thr Leu 120 125 Asn Phe Gly Glu Ser Leu Lys Gly Ile Ser Gln Gly Cys Asn Ile Ile 135 140 Glu Ala Glu Ile Gln Thr Asp Lys Gly Ala Trp Thr Phe Asn Phe Asp 25 155 Lys

(2) INFORMATION FOR SEQ ID NO:94:

30

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 337 amino acids
 - (B) TYPE: amino acid
 - (D) TOPOLOGY: linear

- (ii) MOLECULE TYPE: protein
- (iii) HYPOTHETICAL: YES
- 40 (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Helicobacter pylori
 - (ix) FEATURE:
 - (A) NAME/KEY: misc_feature
- 45 (B) LOCATION 1...337
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:94:
- Met Ile Arg Leu Lys Gly Leu Asn Lys Thr Leu Lys Thr Ser Leu Leu
 50 1 5 10 15

 Ala Gly Val Leu Leu Gly Ala Thr Ala Pro Leu Met Ala Lys Pro Leu
 20 25 30

 Leu Ser Asp Glu Asp Leu Leu Lys Arg Val Lys Leu His Asn Ile Lys
 35 40 45

 55 Glu Asp Thr Leu Thr Ser Cys Asn Ala Lys Val Asp Gly Ser Gln Tyr

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		50					55					60				
		Asn	Ser	Gly	Trp	Asn 70	Leu	Ser	Lys	Glu	Phe 75	Pro	Gln	Glu	Tyr	Arg 80
_	65 Glu	Lys	Ile	Phe			Val	Glu	Glu	Glu 90		His	Lys	Gln	Ala 95	Leu
5	Asn	Leu	Ile	Asn	85 Lys	Glu	Asp	Thr			Lys	Glu	Glu	Leu		Lys
				100					105	_		_	_	110		5 1
	-		115					120					125	Gln		
10	Met	Ala 130	Phe	Glu	Met	Lys	Glu 135	His	Ser	Lys	Glu	Phe 140	Pro	Asn	Lys	Lys
	Gln		Gln	Thr	Met		Glu	Asn	Ala	Phe		Asn	Gly	Ala	Glu	
	145	- 1 -	N	7	Term	150	Glu	7 ~~	Dhe	Glv	155	Tla	Ser	Arg	Glu	160 Asn
15			_		165					170					175	
		_		180					185					Gly 190		
	Leu	Ala	Phe 195	Gly	Glu	Arg	Ser	Tyr 200	Ile	Arg	Gln	Tyr	Lys 205	Lys	Asp	Phe
20	Glu	Glu 210	Ser	Thr	Tyr	Asp	Thr 215	Arg	Gln	Thr	Leu	Ser 220	Ala	Met	Ala	Asn
			Gly	Glu	Asn			Lys	Ile	Thr			Lys	Pro	Lys	
	225	.	TT -	C	C - ~	230	λαπ	Tla	Lare	Pro	235	Met	Ser	Asn	Thr	240 Glu
25					245					250					255	
				260					265					Tyr 270		
	Gly	Cys	Asn 275	Met	Glu	Ile	Asp	Gly 280	Ser	Lys	Tyr	Pro	Ile 285	His	Lys	Asp
30	Trp	Gly 290	Phe	Phe	Gly	Lys	Ala 295	Lys	Val	Pro	Glu	Thr 300	Trp	Arg	Asn	Lys
	Ile		Glu	Cys	Ile	Lys	Asn	Lys	Val	Lys	Ser	Tyr	Asp	Asn	Thr	Thr
	305	_				310					315				_	320
35	Ala	Glu	Ile	Gly	Ile 325	Val	Trp	Lys	Lys	Asn 330	Thr	Tyr	Ser	Ile	Ser	HIS
	His															
	(2)	INF	ORMA'	rion	FOR	SEQ	ID I	NO : 9	5:							
40		(;	SE	OUEN	CE C	HARA	CTER	ISTI	CS :							•
		((2	A) LI	ENGT	H: 4	16 aı	mino		ds						
			-	B) T							•					
45		(ii) MO	LECU	LE T	YPE:	pro	tein								
		•	,				_									
			,	POTH												
50		(vi	•	IGIN A) O				icob	acte	r py	lori					
			,													

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(ix) FEATURE:

(A) NAME/KEY: misc_feature(B) LOCATION 1...416

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:95:

5	Met 1	Lys	Lys	Leu	Val 5	Phe	Ser	Met	Leu	Leu 10	Cys	Cys	Lys	Ser	Val 15	Phe
	Ala	Glu	Gly	Glu 20	Thr	Pro	Leu	Ile	Val 25	Asn	Asp	Pro	Glu	Thr 30	His	Val
			Ala 35				-	40			_		45	_		-
10	Glu	Glu 50	Ile	Ile	Ser	Lys	Ala 55	Gln	Ala	Gln	Val	Asn 60	Gln	Leu	Gln	Lys
	65		Asn			70					75					80
15			Leu		85					90					95	
			_Ser	100		_		_	105			_		110		
20		_	Asn 115					120				_	125			
20	-	130	Trp				135					140				
	145	-	Arg			150				_	155	-				160
25			Asn	_	165					170					175	_
	_	_	Met	180					185					190		
30	-		195 Leu		_			200		-			205			
		210	Leu				215					220				
	225 Asp	Lys	Leu	Asp	Pro	230 Phe	Leu	Lys	Arg	Leu	235 Asp	Val	Leu	Gln	Thr	240 Glu
35	Phe	Gly	Val	Thr	245 Asp	Pro	Thr	Ala	Asn	250 His	Asn	Lys	Gln	Gly	255 Ile	His
	Tyr	Cys	Thr	260 Glu	Asn	Lys	Glu	Thr	265 Gly	Lys	Cys	Asp	Pro	270 Ile	Lys	Asn
40	Val	Phe	275 Arg	Thr	Thr	Arg	Leu	280 Asp	Asn	Glu	Leu	Glu	285 Gln	Glu	Ile	Gln
		290 Leu	Thr		_							_	_		Gln	
45	305 Gln	Ala	Tyr		Asn	310 Phe				Ile						320 Lys
45	Tyr	Leu	Lys		325 Ile	Thr	Asn	Gln		330 Leu	Phe	Leu	Asn		335 Thr	Met
	Ala	Met	Gln	340 Ser	Glu	Ile	Met		345 Asp	Asp	Tyr	Phe		350 Gln	Asn	Asn
50	Asp		355 Phe	Gly	Glu	Lys		360 Asn	His	Ile	Asp		365 Gln	Leu	Thr	Gln
		370 Arg	Ile	Asn	Glu		375 Glu	Arg	Ala	Arg		380 Tyr	Phe	Gln	Asn	Pro 400
55	385 Asn	Val	Lys	Phe	Asp	390 Gln	Phe	Gly	Phe	Pro	395 Ile	Phe	Ser	Ile	Trp	
					x U J					= = 0						

(2) INFORMATION FOR SEQ ID NO:96:

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5		(i)	(A (B		NGTH PE:	: 37 amin	6 am			.s						
10				ECUL				ein								
	(iii)	HYP	OTHE	TICA	L: Y	ES									
15		(vi)	ORI	GINA A) OR	L SC	URCE SM:	: Heli	coba	cter	pyl	ori.					
13		(ix)	(P	ATURE A) NA B) LC	ME/F			fea 376	iture	:					,	
20		(xi)	SEÇ	OUENC	E DE	SCR	PTIC	ON: S	EQ I	D NC	96:	:				
	1				5					10				Gly	T 2	
25	Thr			20					25					Lys 30		
20			35					40					45	ГÀг		
	Leu	Gln 50	Asp	Tyr	Thr	Phe	Thr 55	Gln	Asn	Pro	Gln	Pro 60	Thr	Asn	Thr	Glu
30	6 5	Ser				70	Ile				75			Leu		80
	Ala				85					90				Lys	22	
35				100					105					Pro 110		
			115					120					125	Ile		
		130					135					140		Phe		
40	145	Asn				150					155			Glu		TO
					165					170					1/5	
45				180			_		185					Ile 190		
			195					200					205			
		210					215					220		Tyr		
50		Asn	Lys	Met	Gly	Glu 230		Arg	Leu	Asp	Ile 235	Val	Trp	Ser	Arg	11e
	225 Ile	Thr	Pro	His	Gly 245	Ile	Asn	Ile	Met	Leu 250	Thr		Ala	Lys	Gly 255	Ala
55	Asp	Ile	Lys	Gly 260	Tyr	Asn	Gly	Leu	Val 265	Gly	Glu	Leu	Ile	Glu 270	Arg	Ası

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Phe Gln Arg Tyr Gly Val Pro Leu Leu Ser Thr Leu Thr Asn Gly 280 Leu Leu Ile Gly Ile Thr Ser Ala Leu Asn Asn Arg Gly Asn Lys Glu 295 300 Glu Val Thr Asn Phe Phe Gly Asp Tyr Leu Leu Leu Gln Leu Met Arg 310 315 Gln Ser Gly Met Gly Ile Asn Gln Val Val Asn Gln Ile Leu Arg Asp 325 330 Lys Ser Lys Ile Ala Pro Ile Val Val Ile Arg Glu Gly Ser Arg Val 10 345 Phe Ile Ser Pro Asn Thr Asp Ile Phe Phe Pro Ile Pro Arg Glu Asn 360 Glu Val Ile Ala Glu Phe Leu Lys 370 15 (2) INFORMATION FOR SEQ ID NO:97: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 916 amino acids (B) TYPE: amino acid 20 (D) TOPOLOGY: linear (ii) MOLECULE TYPE: protein 25 (iii) HYPOTHETICAL: YES (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori 30 (ix) FEATURE: (A) NAME/KEY: misc feature (B) LOCATION 1...916 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:97: 35 Val Asp Leu Arg Ile Gln Ser Lys Glu Val Ser His Asn Leu Lys Glu 10 Leu Ser Lys Thr Leu Ile Ser Tyr Pro Phe Glu Lys His Val Glu Ala 25 40 Leu Gly Glu Gln Cys Ser Asn Phe Val Ser Ile Pro Ile Asn Asn Asp 40 Asp Tyr Ser Asn Ile Cys Thr Phe Val Ser Asp Phe Ile Asn Leu Ile Ala Ser Tyr Asn Leu Leu Glu Ser Phe Leu Asp Phe Tyr Lys Asp Lys 45 70 75 Leu Lys Leu Ser Glu Leu Val Thr Glu Tyr Ala Asn Val Thr Asn Asn 85 90 Leu Leu Phe Lys Lys Leu Ile Lys His Leu Ser Gly Asn Asn Gln Leu 105 Val Lys Asn Phe Tyr Gln Cys Ile Arg Glu Ile Ile Lys Tyr Asn Ala 50 120 Pro Asn Lys Glu Tyr Lys Pro Asn Gln Phe Phe Ile Ile Gly Lys Gly 135 140 Lys Gln Lys Gln Leu Ala Lys Ile Tyr Ser His Leu Lys Glu Leu Ser 55 150 · 155

					165					170	Asp				175	
				180					185		Asp			190		
5			195					200			Glu		205			
		210	Asn				215				Asn	220				
10	225	Glu				230					Phe 235					240
10	Ser				245					250	Lys				255	
				260					265		Leu			270		
15			275	Lys				280			Ala		285			
	_	290	Ile				295				Tyr	300				
20	305	Leu				310					Arg 315					320
20	Glu				325					330	Ile				335	
				340					345		Ile			350		
25	_		355					360			Gln		365			
		370					375				Pro	380				
30	385					390					His 395					400
					405					410					415	
				420					425		Leu			430		
35	_		435					440					445			Asn
		450					455					460				Ile
40	465					470					475					Phe 480
					485					490	1				495	Lys
	•			500					505					510		Ser
45			515					520)				525)		Asp
		530					535					540	1			Tyr
50	545					550					555					Lys 560
					565					570)				5/5	
				580)				585	5				590	,	Glu
55	His	Glu	Ile	Lys	Leu	Glu	Val	Туг	Asp	Cys	a Arg	Lys	: Sei	Hls	ASP	His

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			595					600					605			
	Asn	Glu	Pro	Ile	Ile	Leu	Ser	Gln	Gln	Ser	Thr	Gly	Phe	Gln	Trp	Ala
		610					615					620				
	Phe	Asn	Phe	Met	Phe	Gly	Phe	Leu	Tyr	Asn	Val	Gly	Ser	His	Phe	Ser
5	625					630					635					640
	Phe	Asn	His	Asn	Ile	Ile	Tyr	Val	Met	Asp	Glu	Pro	Ala	Thr	His	Leu
					645					650					655	
	Ser	Val	Pro	Ala	Arg	Ļys	Glu	Phe	Arg	Lys	Phe	Leu	Lys	Glu	Tyr	Ala
				660					665					670		
10	His	Lys	Asn	His	Val	Thr	Phe	Val	Leu	Ala	Thr	His	Asp	Pro	Phe	Leu
		_	675					680					685			
	Val	Asp	Thr	Asp	His	Leu	Asp	Glu	Ile	Arg	Ile	Val	Glu	Lys	Glu	Thr
		690					695					700				
	Glu	Gly	Ser	Val	Ile	Lys	Asn	His	Phe	Asn	Tyr	Pro	Leu	Asn	Asn	Ala
15	705					710					715					720
	ser	Lys	Asp	Ser	Asp.	-Ala	Leu	Asp	Lys	He	-Lys-	-Arg-	Ser	Leu	Gly	-Va-1
					725					730					735	
	Gly	Gln	His	Val	Phe	His	Asn	Pro	Gln	Lys	His	Arg	Ile	Ile	Phe	Val
				740					745					750		
20	Glu	Gly	Tle	Thr	Asp	Tyr	Cys	Tyr	Leu	Ser	Ala	Phe	Lys	Leu	Tyr	Leu
			755					760					765			
	Arg	Tyr	Lys	Glu	Tyr	Lys	Asp	Asn	Pro	Ile	Pro	Phe	Thr	Phe	Leu	Pro
		770					775					780			•	
	Ile	Ser	Gly	Leu	Lys	Asn	Asp	Ser	Asn	Asp	Met	Lys	Glu	Thr	Ile	Glu
25	785					790					795					800
	Lys	Leu	Cys	Glu	Leu	Asp	Asn	His	Pro	Ile	Val	Leu	Thr	Asp	Asp	Asp
					805					810					815	
	Arg	Lys	Cys	Val	Phe	Asn	Gln	Gln	Ala	Thr	Ser	Glu	Arg	Phe	Lys	Arg
				820					825					830		
30	Ala	Asn	Glu	Glu	Met	His	Asp		Ile	Thr	Ile	Leu		Leu	Ser	Asp
			835					840					845			
	Cys	Asp	Arg	His	Phe	Lys		Ile	Glu	Asp	Cys		Ser	Ala	Asn	Asp
		850					855					860			_	
	Arg	Asn	Lys	Tyr	Ala	Lys	Asn	Lys	Gln	Met		Leu	Ser	Met	Ala	
35	865					870					875				_	880
	Lys	Thr	Arg	Leu		Tyr	Gly	Gly	Glu		Ala	Ile	Glu	Lys		Thr
					885					890				_	895	
	Lys	Arg	Asn		Leu	Lys	Leu	Phe		Trp	Ile	Ala	Trp		Thr	Asn
40				900					905					910		
40	Leu	Ile	-	Asn												
			915													

(2) INFORMATION FOR SEQ ID NO:98:

- 45 (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 176 amino acids
 - (B) TYPE: amino acid
 - (D) TOPOLOGY: linear
- 50 (ii) MOLECULE TYPE: protein
 - (iii) HYPOTHETICAL: YES
 - (vi) ORIGINAL SOURCE:
- 55 (A) ORGANISM: Helicobacter pylori

SMEDOCID -MIO 081833341 1 -

		(ix)		ATURE				_								
				A) NA					ature	2						
5			(1	3) LC	CAT	LON	L	. / 6								
5		(xi)	SEÇ	QUENC	E DE	ESCRI	[PTIC	ON: 5	SEQ I	D NO	98:	:				
	1				5					10				Thr	15	
10	Phe			20					25					Leu 30		
	_		35					40					45	Val		
15	_	50					55					60		Gly		
	65					70					75			Arg		80
					85					90				Asn	95	
20				100					105					Gly 110		
			115					120					125	Val		
25		130					135					140		Tyr		
	145					150					155			Phe		160
	Val	Leu	Ile	Gly	Ser 165	Val	Val	Phe	Phe	170		ile	Tyr	Trp	175	ne.
30	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO : 9	9:					•		
		(i		QUEN												
35			(A) L B) T	YPE:	ami	no a	cid	acı	as						
				D) T												
		(11) MO	LECU	ו יצע	YPE:	pro	rem								
40		(iii) HY	POTH	ETIC	AL:	YES									
		(vi		IGIN (A) O				icob	acte	r py	lori					
45		(ix	(ATUR (A) N (B) L	AME/				atur	e						
50		(xi) SE	QUEN	CE I	ESCR	IPTI	ON:	SEQ	ID N	10:99):				
J U	Met 1	Phe	Lys	. Asn	Ala 5	. Leu	. Asn	ıle	Gln	Asp 10	Phe	. Ser	Phe	Lys	Asn 15	His
	Thr	Ser	Thr	Ala		: Ile	e Gly	Thr	Asn 25		r Ala	Gly	Lys	Ser 30	Thr	Le
55	-1-	. 7.00	The	- Tle	Lai	Gla	, T1e	Arc	. Ser	Ast	Tvr	Asn	Phe	Lys	Ala	Gl

	3	7	35	T] 0	Dro	The same	uie	40	Acn	Va 1	Tla	Pro	45 Gln	Arg	Lare	GI.
		50				•	55					60				
5	Leu 65	Gly	Val	Val	Ser	Asn 70	Leu	Phe	Asn	Tyr	Pro 75	Pro	Gly	Leu	Asn	80 81
	Asn	Asp	Leu	Phe	Lys 85	Phe	Tyr	Gln	Phe	Phe 90	His	Lys	Asn	Cys	Thr 95	Le
	Asp	Leu	Phe	Glu 100		Asn	Leu	Leu	Asn 105	Lys	Thr	Tyr	Glu	His 110	Leu	Se
10	Asp	Gly			Gln	Arg	Leu	Lys 120		Asp	Leu	Ala	Leu 125	Ser	His	Hi
	Pro		115 Leu	Val	Ile	Met	_		Pro	Glu	Thr			Glu	Gln	Ası
		130	T10	7 ~~~	T 011	eo~	135	Lau	Tla	Sar	Len	140	λαπ	Thr	Gla	G1,
15	145	rea	116	Arg	пеп	150	ASII	пеп	116	Ser	155	Arg	ASII	1111	GIII	160
	-Leu-	-Thr-	-Ser-	-I-le-	-I-l-e-		-Thr	His-	Asp	Pro-		-Va-1	-Leu	Asp	Ser	
					165					170					175	
	Glu	Trp	Val	Leu 180	Leu	Leu	Lys	Asn	Gly 185	Asn	Ile	Ala	Gln	Tyr 190	Lys	Pro
20	Leu	Asn	Ser 195	Ile	Leu	Lys	Ser	Val 200	Ala	Lys	Thr	Phe	Asn 205	Phe	Lys	Glı
	Lys	Pro 210	Thr	Thr	Lys	Asp	Leu 215	Leu	Ala	Leu	Leu	Lys 220	Asp	Ile		
25	(2)	INF	ORMA:	rion	FOR	SEQ	ID 1	NO:10	00:							-
		(i)		_		IARA				_						
						H: 40			acio	is						
30			•			OGY:										
			_													
		(ii)	MOI	LECUI	LE T	YPE:	prot	cein						*		
35		(iii)	HYI	POTH	ETIC	AL: Y	YES									
,,		(vi)				OURCI										
			(2	A) OI	RGAN	ISM:	Heli	icoba	acte	py]	lori					
		(ix)		ATURI												
40						CEY:		_	ature	2						
		(v i)				ESCRI			EO 3	או כד) · 1 () (٠.				
				_												
45	Met 1	Tyr	Ala	Ala	His 5	Pro	Iļe	Lys	Pro	Ile 10	Lys	Ala	Pro	Lys	Leu 15	Lys
	Ser	Gln	Phe	Leu 20	Arg	Arg	Val	Phe	Val 25	Gly	Ala	Ser	Ile	Arg 30	Arg	Trp
50	Asn	Asp	Gln 35		Cys	Pro	Leu	Glu 40	Phe	Val	Glu	Leu	Asp	Lys	Gln	Ala
		Lys 50		Met	Ile	Ala	Tyr 55		Leu	Ala	Lys	Asp 60		Lys	Asp	Arg
	Gly		Asp	Leu	Asp			Leu	Leu	Ile		_	Phe	Cys	Phe	
55	65 Dho	T 0	G1	n ~~	7 6	70 37a 1	Len	ጥኮ~	Δεν	Tla	75 Tuc	Dro	Dro	Tle	Phe	80 TV1

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												•				
					85					90					95	
	21-	T 011	Gl n	Gln	Thr	His	Ser	Lvs	Glu		Ala	Ser	Tyr	Val	Ala	Gln
				100					105					TIO		
	Ser	Leu	Gln	Asp	Glu	Ile	Ser	Ala	Tyr	Phe	Ser	Leu	Glu	Glu	Leu	Lys
5			115					120					125			
	Glu	Tyr	Leu	Ser	His	Arg		Gln	Ile	Leu	Glu	Thr	GIn	TIE	Leu	GIU
		130			_		135	T		<i>~</i> 1	Dha	140	Tle	Tle	TVY	His
		Ala	His	Phe	Tyr	Ala 150	Ser	ьys	rrb	GIU	155	ASP	110		-1-	160
10	145	3	Dwo	7 5 77	Mat	Tyr	Glv	Val	Lvs	Glu		Lvs	Asp	Lys	Ile	Asp
10					165					170					T \ 2	
	Lvs	Gln	Leu	His	Asn	Asn	Asp	His	Leu	Phe	Glu	Gly	Leu	Phe	Gly	Glu
	_			180					185					190		
	Lys	Glu	Asp	Leu	Lys	Lys	Leu	Val	Ser	Met	Phe	Gly	Gln	Leu	Arg	Phe
15			195				,	200	-	777	D	~1 ~	205	Sar	17a l	T. 2 11
	Gln		Arg	Trp	Ser	Gln		Pro	Arg	vaı	PIO	220	1111	361	Val	200
		210	mb	7 011	Cve	Val	215 Ala	Tle	Met	Glv	Tvr		Leu	Ser	Phe	Asp
	G1y 225	HIS	Thr	пеп	Cys	230	MIG			1	235					240
20	Len	Lvs	Ala	Cvs	Lys	Ser	Met	Arg	Ile	Asn	His	Phe	Leu	Gly	Gly	Leu
20					245					250					255	
	Phe	His	Asp	Leu	Pro	Glu	Ile	Leu	Thr	Arg	Asp	Ile	Ile	Thr	Pro	TIE
				260			_		265	a	т1 о	Two	G111	270	Glu	Lvs
~-	Lys	Gln			Ala	Gly	Leu	280	HIS	Cys	116	пуs	285			
25	T	<i>C</i> 3.11	275	Gln	Δen	Lys	Va1		Ser	Phe	Val	Ser		Gly	Val	Gln
	_	290					295					300				
	Glu	Asp	Leu	Lys	Tyr	Phe	Thr	Glu	Asn	Glu	Phe	Lys	Asn	Arg	Tyr	Lys
	205					310					315					320
30	Asp	Lys	Ser	His		Ile	Val	Phe	Thr	Lys	Asp	Ala	GIU	GIU	335	PHE
				_	325		~ 1	The exe	T.011	330		Cvs	Glv	Glu		Leu
	Thr	Leu	. Tyr	Asn 340		Asp	GIU	TYL	345	. Gry	***	- 1	1	350		
	Lare	Va1	Cvs	Asp	His	Leu	Ser	Ala			Glu	Ala	Gln	Ile	Ser	Leu
35	_		355					360					365			
,,,	Ser	His	Gly	Ile	Ser	Ser	Туг	Asp	Leu	ılle	Gln	Gly	Ala	Lys	Asn	Leu
		370	ı				375					380				
	Leu	Glu	Lev	Arg	Ser			GIU	Lev	ı Lev	395	Leu	Asp	реч	, Gry	Lys 400
40	385				Dhe	390					393					
40	Leu	Pne	Arg	ASE	405	Lys :	,									
					40-	•										•
	(2)	INE	ORMA	OIT	FOF	SEC	ID Q	NO: 1	.01:							
	, - ,															
45		i)	L) SI	EQUE	ICE (CHARA	CTE	RISTI	CS:							
						TH: 3			ac	Las						
						: ami LOGY:				•						
				נ (ם)	COPO	LOGI.	. 111	ıcaı								
50		(1)	i) Mo	OLECT	πE :	TYPE:	pro	oteir	ı							
50		\ .	_,				-									
		(iii	i) H	YPOTI	HETI	CAL:	YES									
		(v:	i) O	RIGII	JAL	SOUR	CE:	141			.1 ~	ł				
55				(A)	ORGA	MEIN	: не.	1100	Jact	=r b	ATOL:	L				

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(ix) FEATURE: (A) NAME/KEY: misc feature (B) LOCATION 1...335 5 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:101: Val Leu Trp Val Leu Tyr Phe Leu Thr Ser Leu Phe Ile Cys Ser Leu Ile Val Leu Trp Ser Lys Lys Ser Met Leu Phe Val Asp Asn Ala Asn 10 25 Lys Ile Gln Gly Phe His His Ala Arg Thr Pro Arg Ala Gly Gly Leu Gly Ile Phe Leu Ser Phe Ala Leu Ala Cys Tyr Leu Glu Pro Phe Glu 15 55 Met Pro Phe Lys Gly Pro Phe Val Phe Leu Gly Leu Ser Leu Val Phe 70 Leu Ser Gly Phe Leu Glu Asp Ile Asn Leu Ser Leu Ser Pro Lys Ile 85 90 Arg Leu Ile Leu Gln Ala Val Gly Val Val Cys Ile Ile Ser Ser Thr 20 105 110 Pro Leu Val Val Ser Asp Phe Ser Pro Leu Phe Ser Leu Pro Tyr Phe 120 Ile Ala Phe Leu Phe Ala Ile Phe Met Leu Val Gly Ile Ser Asn Ala 25 135 140 Ile Asn Ile Ile Asp Gly Phe Asn Gly Leu Ala Ser Gly Ile Cys Ala 155 150 Ile Ala Leu Leu Val Ile His Tyr Ile Asp Pro Ser Ser Leu Ser Cys 170 30 Leu Leu Ala Tyr Met Val Leu Gly Phe Met Val Leu Asn Phe Pro Ser 180 185 Gly Lys Ile Phe Leu Gly Asp Gly Gly Ala Tyr Phe Leu Gly Leu Val 200 Cys Gly Ile Ser Leu Leu His Leu Ser Leu Glu Gln Lys Ile Ser Val 35 215 220 Phe Phe Gly Leu Asn Leu Met Leu Tyr Pro Val Ile Glu Val Leu Phe 230 235 Ser Ile Leu Arg Arg Lys Ile Lys Arg Gln Lys Ala Thr Met Pro Asp 250 245 40 Asn Leu His Leu His Thr Leu Leu Phe Lys Phe Leu Gln Gln Arg Ser 265 Phe Asn Tyr Pro Asn Pro Leu Cys Ala Phe Ile Leu Ile Leu Cys Asn 280 Leu Pro Phe Ile Leu Ile Ser Val Leu Phe Arg Leu Asp Ala Tyr Ala 45 295 300 Leu Ile Val Ile Ser Leu Val Phe Ile Ala Cys Tyr Leu Ile Gly Tyr 315 Ala Tyr Leu Asn Arg Gln Val Cys Ala Leu Glu Lys Arg Ala Phe 330 50 (2) INFORMATION FOR SEQ ID NO:102: (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 96 amino acids

(B) TYPE: amino acid

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		(D) TO	POLOGY:	linear					
	(ii)	MOLECUI	LE TYPE:	protein					
5	(iii)	нүротні	ETICAL: Y	(ES					
	(vi)		AL SOURCE RGANISM:		acter pyl	ori			
10	(ix)		E: AME/KEY: OCATION		ature				
15	(xi)) SEQUEN	CE DESCR	IPTION: S	SEQ ID NO	0:102:			
13	Met Lys	Lys Val	Ile Val	Ala Leu	Gly Val	Leu Ala	Phe Al	a Asn 15	Val
	Leu Met	Ala Thr	Asp Val	Lys Ala	Leu Val 25	Lys Gly	Cys Al		Cys
20	His Gly		Phe Glu	Lys Lys	Ala Leu	Gly Lys	Ser Ly 45	s Ile	Val
	Asn Met		Glu Lys	Glu Ile 55	Glu Glu	Asp Leu 60	Met Al	.a Phe	Lys
25		Ala Asn	Lys Asn 70	Pro Val	Met Thr	Ala Gln 75	Ala Ly	/s Lys	Let 80
23	Ser Asp	Glu Asp	Ile Lys 85	Ala Leu	Ala Lys 90	Tyr Ile	Pro Th	nr Leu 95	Lys
30	-) SEQUENC (A) Li (B) T	FOR SEQ CE CHARAGENGTH: 1: YPE: amin	CTERISTION SE AMINO ACID	CS:		,		
35	(ii) MOLECU	LE TYPE:	protein					
	(iii) НҮРОТН	ETICAL:	YES					
40	(vi		AL SOURCE		acter py	lori			
45	(ix		E: AME/KEY: OCATION		ature				
	(xi) SEQUEN	CE DESCR	IPTION:	SEQ ID N	0:103:			
50	1		Asn Asn 5		10			15	
		20	Glu Lys Ala Ser		25		3	0	
55		35	Phe Gly	40			45		

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		50					55					60				
	Val 65	Glu	Leu	Ile	Ala	Pro 70	Phe	Leu	Asp	Thr	Glu 75	Glu	Tyr	Gly	Ile	Pho 80
5	Arg	Glu	Asp	Glu	His 85	Glu	Pro	Leu	Val	Ile 90	Ser	Val	Ile	Lys	Lys 95	Gl
	Lys	Thr	Arg	Tyr 100	Phe	Leu	Asn	Gln	Thr 105	Ser	Leu	Ser	Lys	Asn 110	Thr	Le
	Lys	Ala	Leu 115	Leu	Lys	Gly	Leu	Ile 120	Lys	Arg	Leu	Ser	Asn 125	Asp	Arg	Phe
10	Ser	Gln 130	Asn	Glu	Leu	Asn	Asp 135	Ile	Leu	Met	Leu	Ser 140	Leu	Leu	Asp	Gl
	Tyr 145	Ile	Gln	Asn	Lys	Asn 150	Arg	Arg	Leu	Ala	Pro 155	Phe				
15	(2)	INFO	ORMA'	rion	FOR	SEQ	ID I	NO:1	04:							
			1	··						-		-	•			-
		(i)				HARA(4.0						
						H: 1: ami:			acı	15						
20			-			OGY:										
		(ii)	MOI	LECUI	LE T	YPE:	prot	tein								
25		(iii)	HYI	POTH	ETIC	AL: Y	ŒS									
		(vi)	OR:	IGINA	AL S	OURCI	Ξ:									
			(2	A) OF	RGAN	ISM:	Hel:	icoba	acte	r py:	lori					
		(= ==)	विव	ATURI	7.											
30		(TX)				KEY:	misc	: fea	ature	2						
						ON I										
		(x1)	SEÇ	QUENC	E DE	ESCRI	PTIC	ON: S	SEQ 1	ED NO):104	ł:				
35	Val 1	Met	Leu	Met	Ala 5	Ile	Phe	Thr	Pro	Tyr 10	Ile	Leu	Ile	Leu	Lys 15	Met
		Lys	Lys	Ser 20	Met	Ser	Leu	Phe	Ala 25		Met	Gly	Leu	Glu 30	Gln	Ile
10	Phe	Cys	Asn 35	Arg	Asp	Ile	Lys	Asp 40	Leu	Asn	Asp	Phe	Val 45	Phe	Gly	Ile
	Glu	Val 50	Gly	Leu	Asp	Ser	Asn 55	Ala	Arg	Lys	Asn	Arg 60	Ser	Arg	Lys	Ala
	Met	Glu	Asn	His	Leu		Gly	Leu	Phe	Val	Gln	Ala	Gln	Leu	Asn	
15	65	~1	~ 1	••- 1	•	70	•	~ 1	51 .	~7	75		_	~ 1		80
15	гÀг	Glu	GIN	vai	Asp 85	iie	Arg	GIU	Pne	90	Asp	Leu	Arg	GIN	95	Pne
	Gly	Asn	Asp	Thr 100		Lys	Phe	Asp	Phe 105		Ile	Phe	Ser	Lys 110		Lys
	Thr	Tyr		His	Arg	Ser										
0			115													
	(2)	INFO	RMAT	NOI	FOR	SEQ	ID N	10:10	5:							
		(3)	SEC	TENC	E CH	מפמר	ו פשיי	STIC	٠c .							

(A) LENGTH: 355 amino acids

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(B) TYPE: amino acid
               (D) TOPOLOGY: linear
         (ii) MOLECULE TYPE: protein
5
        (iii) HYPOTHETICAL: YES
         (vi) ORIGINAL SOURCE:
               (A) ORGANISM: Helicobacter pylori
10
         (ix) FEATURE:
               (A) NAME/KEY: misc_feature
               (B) LOCATION 1...355
         (xi) SEQUENCE DESCRIPTION: SEQ ID NO:105:
15
     Met Asn Ile Lys Ile Leu Lys Ile Leu Val Gly Gly Leu Phe Phe Leu
                                         10
     Ser Leu Asn Ala His Leu Trp Gly Lys Gln Asp Asn Ser Phe Leu Gly
                                     25
20
     Ile Gly Glu Arg Ala Tyr Lys Ser Gly Asn Tyr Ser Lys Ala Ala Ser
     Tyr Phe Lys Lys Ala Cys Asn Asp Gly Val Ser Glu Gly Cys Thr Gln
     Leu Gly Ile Ile Tyr Glu Asn Gly Gln Gly Thr Arg Ile Asp Tyr Lys
25
     Lys Ala Leu Glu Tyr Tyr Lys Thr Ala Cys Gln Ala Asp Asp Arg Glu
                                         90
     Gly Cys Phe Gly Leu Gly Gly Leu Tyr Asp Glu Gly Leu Gly Thr Ala
                                    105
30
     Gln Asn Tyr Gln Glu Ala Ile Asp Ala Tyr Ala Lys Ala Cys Val Leu
                                 120
     Lys His Pro Glu Ser Cys Tyr Asn Leu Gly Ile Ile Tyr Asp Arg Lys
                                                 140
                            135
     Ile Lys Gly Asn Ala Ala Gln Ala Val Thr Tyr Tyr Gln Lys Ser Cys
35
                                             155
                         150
     Asn Phe Asp Met Ala Lys Gly Cys Tyr Ile Leu Gly Thr Ala Tyr Glu
                                        170
     Lys Gly Phe Leu Glu Val Lys Gln Ser Asn His Lys Ala Val Ile Tyr
                                     185
40
                 180
     Tyr Leu Lys Ala Cys Arg Leu Asn Glu Gly Gln Ala Cys Arg Ala Leu
                                 200
     Gly Ser Leu Phe Glu Asn Gly Asp Ala Gly Leu Asp Glu Asp Phe Glu
                             215
     Val Ala Phe Asp Tyr Leu Gln Lys Ala Cys Ala Leu Asn Asn Ser Gly
45
                         230
                                             235
     Gly Cys Ala Ser Leu Gly Ser Met Tyr Met Leu Gly Arg Tyr Val Lys
                                         250
                     245
     Lys Asp Pro Gln Lys Ala Phe Asn Tyr Phe Lys Gln Ala Cys Asp Met
50
                                     265
                 260
     Gly Ser Ala Val Ser Cys Ser Arg Met Gly Phe Met Tyr Ser Gln Gly
                                 280
     Asp Thr Val Ser Lys Asp Leu Arg Lys Ala Leu Asp Asn Tyr Glu Arg
                             295
                                                 300
     Gly Cys Asp Met Gly Asp Glu Val Gly Cys Phe Ala Leu Ala Gly Met
55
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305 310 315 Tyr Tyr Asn Met Lys Asp Lys Glu Asn Ala Ile Met Ile Tyr Asp Lys 325 330 Gly Cys Lys Leu Gly Met Lys Gln Ala Cys Glu Asn Leu Thr Lys Leu Arg Gly Tyr (2) INFORMATION FOR SEQ ID NO:106: 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 193 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: protein (iii) HYPOTHETICAL: YES 20 (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori (ix) FEATURE: (A) NAME/KEY: misc_feature 25 (B) LOCATION 1...193 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:106: Met Lys Glu Lys Asn Phe Trp Pro Leu Gly Ile Met Ser Val Leu Ile 30 Phe Gly Leu Gly Ile Val Val Phe Leu Val Val Phe Ala Leu Lys Asn 25 Ser Pro Lys Asn Asp Leu Val Tyr Phe Lys Gly His Asn Glu Val Asp 40 Leu Asn Phe Asn Ala Met Leu Lys Thr Tyr Glu Asn Phe Lys Ser Asn 35 Tyr Arg Phe Ser Val Gly Leu Lys Pro Leu Thr Glu Ser Pro Lys Thr 70 75 Pro Ile Leu Pro Tyr Phe Ser Lys Gly Thr His Gly Asp Lys Lys Ile 40 90 85 Gln Glu Asn Leu Leu Asn Asn Ala Leu Ile Leu Glu Lys Ser Asn Thr Leu Tyr Ala Gln Leu Gln Pro Leu Lys Pro Ala Leu Asp Ser Pro Asn 120 Ile Gln Val Tyr Leu Ala Phe Tyr Pro Ser Gln Ser Gln Pro Arg Leu 45 135 Leu Gly Thr Leu Asp Cys Lys Asn Ala Cys Glu Pro Leu Lys Phe Asp 150 155 Leu Leu Glu Gly Asp Lys Val Gly Arg Tyr Lys Ile Leu Phe Lys Phe 50 170 Val Phe Lys Asn Lys Glu Glu Leu Ile Leu Glu Gln Leu Ala Phe Phe 185

Lys

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(2) INFORMATION FOR SEQ ID NO:107:
          (i) SEQUENCE CHARACTERISTICS:
               (A) LENGTH: 289 amino acids
               (B) TYPE: amino acid
5
               (D) TOPOLOGY: linear
         (ii) MOLECULE TYPE: protein
        (iii) HYPOTHETICAL: YES
10
         (vi) ORIGINAL SOURCE:
               (A) ORGANISM: Helicobacter pylori
         (ix) FEATURE:
15
               (A) NAME/KEY: misc_feature
               (B) LOCATION 1...289
         (xi) SEQUENCE DESCRIPTION: SEQ ID NO:107:
20
    Leu Gly Ile Asn Met Cys Ser Lys Lys Ile Arg Asn Leu Ile Leu Cys
     Phe Gly Phe Ile Leu Ser Leu Cys Ala Glu Glu Asn Ile Thr Lys Glu
                                     25
     Asn Met Thr Glu Thr Asn Thr Thr Glu Glu Asn Thr Pro Lys Asp Ala
25
     Pro Ile Leu Leu Glu Glu Lys Arg Ala Gln Thr Leu Glu Leu Lys Glu
                             55
     Glu Asn Glu Val Ala Lys Lys Ile Asp Glu Lys Ser Leu Leu Glu Glu
                                             75
30
     Ile His Lys Lys Lys Arg Gln Leu Tyr Met Leu Lys Gly Glu Leu His
                                        90
                    85
     Glu Lys Asn Glu Ser Ile Leu Phe Gln Gln Met Ala Lys Asn Lys Ser
                                     105
     Gly Phe Phe Ile Gly Val Ile Leu Gly Asp Ile Gly Ile Asn Ala Asn
35
                                                     125
                                 120
     Pro Tyr Glu Lys Phe Glu Leu Leu Ser Asn Ile Gln Ala Ser Pro Leu
                                                 140
                             135
     Leu Tyr Gly Leu Arg Ser Gly Tyr Gln Lys Tyr Phe Ala Asn Gly Ile
                                             155
40
     Ser Ala Leu Arg Phe Tyr Gly Glu Tyr Leu Gly Gly Ala Met Lys Gly
     Phe Lys Ser Asp Ser Leu Ala Ser Tyr Gln Thr Ala Ser Leu Asn Ile
                                     185
     Asp Leu Leu Met Asp Lys Pro Ile Asp Lys Glu Lys Arg Phe Ala Leu
45
                                                      205
                                 200
     Gly Ile Phe Gly Gly Val Gly Val Gly Trp Asn Gly Met Tyr Gln Asn
                             215
     Leu Lys Glu Ile Arg Gly Tyr Ser Gln Pro Asn Ala Phe Gly Leu Val
50
                                              235
                         230
     Leu Asn Leu Gly Val Ser Met Thr Leu Asn Leu Lys His Arg Phe Glu
```

250

Leu Ala Leu Lys Met Pro Pro Leu Lys Glu Thr Ser Gln Thr Phe Leu 260 265 270

Tyr Tyr Phe Lys Ser Thr Asn Ile Tyr Tyr Ile Ser Tyr Asn Tyr Leu

245

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275 280 285 Leu 5 (2) INFORMATION FOR SEQ ID NO:108: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 668 amino acids (B) TYPE: amino acid 10 (D) TOPOLOGY: linear (ii) MOLECULE TYPE: protein (iii) HYPOTHETICAL: YES 15 (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori (ix) FEATURE: 20 (A) NAME/KEY: misc_feature (B) LOCATION 1...668 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:108: 25 Met Arg Lys Leu Phe Ile Pro Leu Leu Phe Ser Ala Leu Glu Ala Asn Glu Lys Asn Gly Phe Phe Ile Glu Ala Gly Phe Glu Thr Gly Leu 25 Leu Glu Gly Thr Gln Thr Gln Glu Lys Arg His Thr Thr Thr Lys Asn 30 40 Thr Tyr Ala Thr Tyr Asn Tyr Leu Pro Thr Asp Thr Ile Leu Lys Arg 55 Ala Ala Asn Leu Phe Thr Asn Ala Glu Ala Ile Ser Lys Leu Lys Phe 70 75 35 Ser Ser Leu Ser Pro Val Arg Val Leu Tyr Met Tyr Asn Gly Gln Leu 85 90 Thr Ile Glu Asn Phe Leu Pro Tyr Asn Leu Asn Asn Val Lys Leu Ser 105 Phe Thr Asp Ala Gln Gly Asn Thr Ile Asp Leu Gly Val Ile Glu Thr 40 120 Ile Pro Lys His Ser Lys Ile Val Leu Pro Gly Glu Ala Phe Asp Ser 135 Leu Lys Glu Ala Phe Asp Lys Ile Asp Pro Tyr Thr Leu Phe Leu Pro 150 155 45 Lys Phe Glu Ala Thr Ser Thr Ser Ile Ser Asp Thr Asn Thr Gln Arg 165 170 Val Phe Glu Thr Leu Asn Asn Ile Lys Thr Asn Leu Ile Met Lys Tyr 185 Ser Asn Glu Asn Pro Asn Asn Phe Asn Thr Cys Pro Tyr Asn Asn Asn 50 200 Gly Asn Thr Lys Asn Asp Cys Trp Gln Asn Phe Thr Pro Gln Thr Ala

215

230

55

Glu Glu Phe Thr Asn Leu Met Leu Asn Met Ile Ala Val Leu Asp Ser

Gln Ser Trp Gly Asp Ala Ile Leu Asn Ala Pro Phe Glu Phe Thr Asn

220

															255	
					245	_		D	D-4-	250	Tira	Cvc	17-1	λen		Glv
				260		Asp			265					270		
5	Val	Asn	Gly 275	Arg	Val	Asp	Thr	Lys 280	Val	Asp	Gln	Gln	Tyr 285	Ile	Leu	Asn
J	Lys		Gly	Ile	Ile	Asn	Asn 295	Phe	Arg	Lys	Lys	Ile 300	Glu	Ile	Asp	Ala
		290 Val	Leu	Lys	Asn	Ser		Val	Val	Gly	Leu 315	Ala	Asn	Gly	Tyr	Gly 320
10	305		~1	~1	/The ex-	310 Gly	ጥኮሎ	T.e11	Glv	Val		Δla	Tvr	Ala	Leu	Asp
10					325					330					335	
				340		Gly			345					350		
15			355			His		360					365			
13	Asn	Gly	Asn	Met	Thr	Tyr	Gln 375	Arg	Val	Pro	Val	Thr 380	Lys	Asp	Gly	Gln
	V=1		Lvs	Asp	Ser	Asn		Lys	Pro	Lys	Asp	Ser	Asp	Gly	Leu	Pro
	385					390					395					400
20	Tvr	Asn	Val	Cys	Ser	Leu	Tyr	Gly	Gly	Ser	Asn	Gln	Pro	Ala	Phe	Pro
					405					410					415	
	Ser	Asn	Tyr	Pro	Asn	Ser	Ile	Tyr	His	Asn	Cys	Ala	Asp	Val	Pro	Ala
				420					425		_		_	430		61 -
	Gly	Phe	Leu	Gly	Val	Thr	Ala		Val	Trp	Gln	Gln		TTE	Asn	GIN
25			435			_	_	440		•	a 1	0	445	Thr	λen	Titr
		450				Asn	455					460				
	Asn	Leu	Asn	Ala	Ser	Leu	Asn	Thr	Gln	Asp		Ala	Asn	Ser	Met	480
	465		_		_	470	- 1 -	** 7	m\	0	475	37-1	Thr	Aen	His	
30					485	Thr				490					495	
				500		Gln			505					510		
35			515			Gln		520					525			
	Tyr	Tyr	Gly	Ile	Ile	Lys	Tyr	Asn	Tyr	Ala	Lys	Ala	Val	Asn	GIn	Lys
		530					535					540		+	7	Dho
		Gln	Gln	Leu	Ser	Tyr 550	Gly	Gly	Gly	Ile	Asp 555	Leu	Leu	гел	Asp	Phe 560
40	545	ሞኮሎ	Thr	ጥኒታን	Ser	Asn	Lvs	Asn	Ser	Pro		Gly	Ile	Gln	Thr	Lys
40					565					570					5/5	
				580					585					590		Leu
	Tyr	Asn	Ser	Tyr	Tyr	Val	Leu			Val	Lys	Gly	ser	GIY	ASH	Leu
45			595			_		600			•	774 -	605		ጥ ላ ታንግ	Ser
		610					615					620				Ser
	Val	Gly	Ile	Ser	Ile			Ile	Gln	Arg	Lys	Ala	ser	vai	val	Ser
	625					630		_			635		~ T -		. אם	640 Ser
50					645					650	t .			. ст	655	Ser
	His	Phe	Lys	Val 660		Phe	Asn	Туг	Gly 665		Val	Phe	!			

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(i) SEQUENCE CHARACTERISTICS:
                (A) LENGTH: 63 amino acids
                (B) TYPE: amino acid
 5
                (D) TOPOLOGY: linear
          (ii) MOLECULE TYPE: protein
         (iii) HYPOTHETICAL: YES
10
          (vi) ORIGINAL SOURCE:
                (A) ORGANISM: Helicobacter pylori
          (ix) FEATURE:
15
               (A) NAME/KEY: misc_feature
          - (B)-LOGATION-1...63
          (xi) SEQUENCE DESCRIPTION: SEQ ID NO:109:
20
     Met Asn Thr Glu Ile Leu Thr Ile Met Leu Val Val Ser Val Leu Met
                                          10
     Gly Leu Val Gly Leu Ile Ala Phe Leu Trp Gly Val Lys Ser Gly Gln
                                     25
     Phe Asp Asp Glu Lys Arg Met Leu Glu Ser Val Leu Tyr Asp Ser Ala
25
                                 40
     Ser Asp Leu Asn Glu Ala Ile Leu Gln Glu Lys Arg Gln Lys Asn
     (2) INFORMATION FOR SEQ ID NO:110:
30
          (i) SEQUENCE CHARACTERISTICS:
               (A) LENGTH: 406 amino acids
               (B) TYPE: amino acid
               (D) TOPOLOGY: linear
35
        (ii) MOLECULE TYPE: protein
        (iii) HYPOTHETICAL: YES
40
         (vi) ORIGINAL SOURCE:
               (A) ORGANISM: Helicobacter pylori
         (ix) FEATURE:
               (A) NAME/KEY: misc feature
45
               (B) LOCATION 1...406
         (xi) SEQUENCE DESCRIPTION: SEQ ID NO:110:
     Met Val Phe Phe His Lys Lys Ile Ile Leu Asn Phe Ile Tyr Ser Leu
50
     Met Val Ala Phe Leu Phe His Leu Ser Tyr Gly Val Leu Leu Lys Ala
                                     25
     Asp Gly Met Ala Lys Lys Gln Thr Leu Leu Val Gly Glu Arg Leu Val
55
     Trp Asp Lys Leu Thr Leu Leu Gly Phe Leu Glu Lys Asn His Ile Pro
```

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		50					55					60				
		Lys	Leu	Tyr	Tyr	Asn 70	Leu	Ser	Ser	Gln	Asp 75	Lys	Glu	Leu	Ser	Ala 80
_	65 Glu	Ile	Gln	Ser			Thr	Tyr	Tyr	Thr 90	Leu	Arg	Asp	Ala	Asn 95	Asn
5	Thr	Leu	Ile		85 Ala	Leu	Ile	Pro			Gln	Asp	Leu	Gln		His
		Tyr	T	100	<i>C</i> 111	C1.	λεπ	ጥላታም	105 Phe	Leu	Asp	Phe	Ile	110 Pro	Ile	Val
			115					120					125			
10		Thr 130					135					140				
	Tyr	Gln	Asp	Ile	Val		Ala	Thr	Asn	Asp	Pro	Leu	Leu	Ala	Asn	160
	145				_	150	•	a	17 7	D~-0	155 Dho	Luc	7 ~~~	T.e.ii	·Wal	
15		Met			165					170					175	
		Asp		180					185					190		
	Ala	Phe	Gly 195	Gln	Pro	Thr	Ile	Lys 200	Met	Ala	Met	Val	Ser 205	Ser	Arg	Leu
20	His	Gln	Tyr	Tyr	Leu	Phe	Ser 215		Ser	Asn	Gly	Arg 220	Tyr	Tyr	Asp	Ser
	Larg	210 Ala	Gln	Glu	Val	Ala		Phe	Leu	Leu	Glu		Pro	Val	Lys	Tyr
	225					230					235					240
25		Arg			245					250					255	
		Val		260					265					270		
	Ser	Leu	Ile 275	His	Ser	Ala	Ser	Asp 280	Gly	Arg	Val	Gly	Phe 285	Ile	Gly	Val
30	Lys	Ala	Gly	Tyr	Gly	Lys	Val 295		Glu	Ile	His	Leu 300	Asn	Glu	Leu	Arg
		290 Val	Tyr	Ala	His			Ala	Phe	Ala	Asn 315		Leu	Lys	Lys	Gly 320
	305 Ser	Phe	Val	Lys	Lys	310 Gly	Gln	Ile	Ile	Gly		Val	Gly	Ser	Thr	
35					325					330					335	
		Ser		340					345					350		
		Ile	355					360					365			
40		Lys 370	Gln	Arg			375					380				
	Lys 385	Leu	Glu	Glu	Leu	Phe 390	Lys	Thr	His	Ser	Phe 395	Glu	Lys	Asn	Ser	Phe 400
		Leu	Leu	Glu	Gly	Phe										
45	, -				405											
	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:1	11:							

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 296 amino acids

(B) TYPE: amino acid

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

55

(iii) HYPOTHETICAL: YES

CDOCID: JUO 0010222A1 1 -

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```
(vi) ORIGINAL SOURCE:
                (A) ORGANISM: Helicobacter pylori
 5
          (ix) FEATURE:
                (A) NAME/KEY: misc feature
                (B) LOCATION 1...296
10
         (xi) SEQUENCE DESCRIPTION: SEQ ID NO:111:
     Leu Phe Leu Val Lys Lys Ile Gly Val Val Ile Met Ile Leu Val Cys
                                          10
     Phe Leu Ala Cys Ser Gln Glu Ser Phe Ile Lys Met Gln Lys Lys Ala
15
                                     25
     Gin Glu Gin Glu Asn Asp Gly Ser Lys Arg Pro Ser Tyr Val Asp Ser
                                  40
     Asp Tyr Glu Val Phe Ser Glu Thr Ile Phe Leu Gln Asn Met Val Tyr
20
     Gln Pro Ile Glu Glu Arg Asn Ala Phe Phe Gln Leu Thr Lys Asp Glu
                         70
                                              75
     Asp Asn Ser Phe Asn Pro Glu Asn Ser Val Ile Leu Leu Asn Glu Pro
                                         90
     Ser Asp Asn Ser Glu Lys Asn Leu Leu Ser Tyr Pro Asn Asp Pro Asn
25
                                      105
     Asn Asn Glu Asp Asn Ala Asn Asn Ser Gln Lys Asn Pro Phe Leu Tyr
     Lys Pro Lys Arg Lys Thr Lys Asn Pro Lys Leu Ile Glu Tyr Ser Gln
                             135
30
     Gln Asp Phe Tyr Pro Leu Lys Asn Gly Asp Ile Ile Met Ser Lys Glu
                         150
     Gly Asp Gln Trp Leu Ile Glu Ile Gln Ser Lys Ala Leu Lys Arg Phe
                     165
                                         170
     Leu Lys Asp Gln Asn Asp Lys Asp Arg Gln Ile Gln Thr Phe Thr Phe
35
                                     185
     Asn Asp Thr Lys Thr Gln Ile Ala Gln Ile Lys Gly Lys Ile Ser Ser
                                 200
     Tyr Val Tyr Thr Thr Asn Asn Gly Ser Leu Ser Leu Arg Pro Phe Tyr
                             215
                                                 220
     Glu Ser Phe Leu Leu Glu Lys Lys Ser Asp Asn Val Tyr Thr Ile Glu
40
                         230
                                             235
     Asn Lys Ala Leu Asp Thr Met Glu Ile Ser Lys Cys Gln Met Val Leu
                     245
                                         250
     Lys Lys His Ser Thr Asp Lys Leu Asp Ser Gln His Lys Ala Ile Ser
45
                                     265
     Ile Asp Leu Asp Phe Lys Lys Glu Arg Phe Lys Ser Asp Thr Glu Leu
                                 280
                                                     285
     Phe Leu Glu Cys Leu Lys Glu Ser
         290
50
     (2) INFORMATION FOR SEQ ID NO:112:
          (i) SEQUENCE CHARACTERISTICS:
               (A) LENGTH: 248 amino acids
55
               (B) TYPE: amino acid
```

```
(D) TOPOLOGY: linear
         (ii) MOLECULE TYPE: protein
 5
        (iii) HYPOTHETICAL: YES
         (vi) ORIGINAL SOURCE:
               (A) ORGANISM: Helicobacter pylori
10
         (ix) FEATURE:
               (A) NAME/KEY: misc_feature
               (B) LOCATION 1...248
         (xi) SEQUENCE DESCRIPTION: SEQ ID NO:112:
15
     Val Ser Tyr Asp Asn Thr Asp Asp Tyr Tyr Phe Pro Arg Asn Gly Val
                                         10
     Ile Phe Ser Ser Tyr Ala Thr Met Ser Gly Leu Pro Ser Ser Gly Thr
                                     25
     Leu Asn Ser Trp Asn Gly Leu Gly Gly Asn Val Arg Asn Thr Lys Val
20
                                 40
     Tyr Gly Lys Phe Ala Ala Tyr His His Leu Gln Lys Tyr Leu Leu Ile
     Asp Leu Ile Ala Arg Phe Lys Thr Gln Gly Gly Tyr Ile Phe Arg Tyr
25
                                             75
                         70
     Asn Thr Asp Asp Tyr Leu Pro Leu Asn Ser Thr Phe Tyr Met Gly Gly
                     85
     Val Thr Thr Val Arg Gly Phe Arg Asn Gly Ser Ile Thr Pro Lys Asp
                                     105
     Glu Phe Gly Leu Trp Leu Gly Gly Asp Gly Ile Phe Thr Ala Ser Thr
30
                                                     125
     Glu Leu Ser Tyr Gly Val Leu Lys Ala Ala Lys Met Arg Leu Ala Trp
                                                 140
                             135
     Phe Phe Asp Phe Gly Phe Leu Thr Phe Lys Thr Pro Thr Arg Gly Ser
35
                         150
                                             155
     Phe Phe Tyr Asn Ala Pro Thr Thr Ala Asn Phe Lys Asp Tyr Gly
                                         170
                    165
     Val Val Gly Ala Gly Phe Glu Arg Ala Thr Trp Arg Ala Ser Thr Gly
                                     185
     Leu Gln Ile Glu Trp Ile Ser Pro Met Gly Pro Leu Val Leu Ile Phe
40
                                 200
     Pro Ile Ala Phe Phe Asn Gln Trp Gly Asp Gly Asn Gly Lys Lys Cys
                                                 220
                             215
     Lys Gly Leu Cys Phe Asn Pro Asn Met Asn Asp Tyr Thr Gln His Phe
45
                         230 .
                                             235
     Glu Phe Ser Met Gly Thr Arg Phe
                     245
     (2) INFORMATION FOR SEQ ID NO:113:
50
          (i) SEQUENCE CHARACTERISTICS:
               (A) LENGTH: 335 amino acids
               (B) TYPE: amino acid
               (D) TOPOLOGY: linear
```

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```
(ii) MOLECULE TYPE: protein
```

(iii) HYPOTHETICAL: YES

5 (vi) ORIGINAL SOURCE:

(A) ORGANISM: Helicobacter pylori

(ix) FEATURE:

(A) NAME/KEY: misc_feature

10 (B) LOCATION 1...335

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:113:

	Val	Gln	His	Phe	Asn	Phe	Leu	Tyr	Lys	Asp	Ser	Leu	Phe	Ser	Ile	Ala
15	1				5					10					15	
	Leu	Phe	Thr	-Phe	Tle	Tle	Ala	Leu	val	Ile	Leu	Leu	Glu	Gln	Ala	Arg
				20					25					30		
	Ala	Tyr		Thr	Arg	Lys	Arg		Lys	Lys	Phe	Leu	Gln	Lys	Phe	Ala
20			35	_		_		40					45			
20	Gln		Gln	Asn	Ala	Tyr		Ser	Ser	Glu	Asn		Asp	Glu	Leu	Leu
	T	50	7.7 -	T	T1 -	C	55	T	N = 4.	Di-	•	60	_			
	65	HIS	ALA	пув	116	Ser	ser	Leu	Mec	Pne	ьеи 75	АТА	Arg	AIA	TYP	ser 80
		Δla	Asn	Va 1	Glu	Met	Ser	Tle	G311	Tla	-	Tare	G1v	T.011	T.A11	
25	_,	,,,,,			85					90	DCu	Lys	GLY	пси	95	ASII
	Arq	Pro	Leu	Lys		Glu	Glu	Lvs	Ile		Val	Leu	Asp	Leu		Ala
				100	-			•	105					110		
	Lys	Asn	Tyr	Phe	Ser	Val	Gly	Tyr	Leu	Gln	Lys	Thr	Lys	Asp	Thr	Val
			115					120			_		125	-		
30	Lys	Glu	Ile	Leu	Arg	Phe	Ser	Pro	Arg	Asn	Val	Glu	Ala	Leu	Leu	Lys
		130					135					140				
		Leu	His	Ala	Tyr	Glu	Leu	Glu	Lys	Asp		Ser	Lys	Ala	Leu	
	145	_		_	_	150		_			155					160
35	Thr	Leu	GIu	Cys		Glu	Glu	Leu	Glu		Pro	Lys	Ile	Glu		IIe
33	T	7	TT1 520	T 011	165	7 011	Made	***	*	170	01		T	a 1	175	N 1 -
	цуs	ASI	TYL	180	Tyr	Leu	Mec	HIS	185	iie	GIU	ASN	гÀг	190	Asp	AIA
	Δla	Lvs	Tle		Hic	Val	Ser	Live		Sar	T.011	λen	Lau		Lare	Tla
	VIG	בעם	195	пец	1113	VAI	261	200	AIA	Ser	neu	мэр	205	цуз	пуs	110
40	Ala	Leu		His	Leu	Lys	Ser		Asp	Glu	Asn	Leu		Trp	Gln	Glu
		210					215					220				
	Ile	Asp	Thr	Thr	Glu	Arg	Leu	Glu	Asn	Val	Ile	Asp	Leu	Leu	Trp	Asp
	225					230					235	•			_	240
	Met	Asn	Ile	Pro	Ala	Phe	Ile	Leu	Glu	Lys	His	Ala	Leu	Leu	Gln	Asp
45					245					250					255	
	Ile	Ala	Arg	Ser	Gln	Gly	Leu	Leu	Leu	Asp	His	Lys	Pro	Cys	Gln	Ile
				260					265					270		
	Phe	Glu		Glu	Val	Leu	Arg		Leu	Leu	His	Ser		Ile	Lys	Ala
50	_	_	275			_		280					285		_	
50	Ser		Thr	Phe	Glu	Tyr		Cys	Lys	His	Cys		Gln	Ile	Phe	Pro
	Dhe	290	Cor	ui a	7	C	295	17-1	G	M	~1	300	• -	nh -	M	N
	305	GIU	ser	uls	Arg	Cys 310	PLO	vaı	cys	TYT		ьeu	Ala	ьпе	Met	
		17a 1	T.em	T.ve	Tle	Ser	Lve	Lare	Thr	ui.	315	Mot	G) v	V-1	Aen	320
55	1-1	• 41		~ ₇ 3	325	J-1	Jy 3	-ys		330.	nid	MEL	GTÄ	Val	335	•

	(2)	INFO	RMAT	NOI	FOR	SEQ	ID N	0:11	4:							
5		(i)	(B	UENC .) LE .) TY .) TO	NGTH PE:	: 41 amin	.3 am	ino id		ls						
10			MOL					ein								
	(iii)	HYP	OTHE	TICA	L: Y	ES									
15		(vi)	ORI (A	GINA () OR				.coba	cter	pyl	ori.					
		(ix)		TURE A) NA B) LC	WE/k			_	ture	£						
20		(xi)	SEÇ	QUENC	E DE	SCRI	PTIC	ON: 5	EQ I	D NO	:114	l:				
	1				5					10					15	Gly
25	Val	Val	Tyr	Ala 20	Glu	Pro	Asp	Ser	Lys 25	Val	Glu	Ala	Leu	Glu 30	Gly	Arg
23	_		Glu 35	Ser				40	Lys				45			
	Lys	Glu 50	Leu	Lys	Asn	Lys	Glu 55	Leu	Lys	Asn	Lys	Asp 60	Leu	Lys	Asn	Lys
30	65	Glu	Lys			70					75					80
			Gly		85					90					95	
35			Gly	100					105					110		
			Lys 115	-				120					125			
		130	Glu				135					140				
40	_	Asn	Asn	Ala	Thr	Asn 150	Asn	Thr	Leu	Glu	Asp 155	Lys	Val	Val	Gly	Gly 160
	145 Ile	Ser	Leu	Leu	Val		Gly	Ser	Pro	Ile		Leu	Tyr	Gln	Ile	Gln
					165					170					175	
45	Glu	Glu	Gln	180	гуs	ser	Lys	vaı	185	гу	Ala	GIII	ALG	190	nop.	Arg
75			195	Glu				200					205			Ile
		210					215					220				Gln
50		Gln	Gly	Met	Asp			His	Phe	Lys			Leu	Met	Ala	Glu 240
	225 Gly	His	Tyr	Lys	Leu 245	230 Tyr		Asp	Gln	Leu 250	235 Lys		His	Leu	Glu 255	Met
<i></i>	Gln	Glu	Leu		Arg	Asn	Ile	Leu	Leu 265	Thr		Val	Asp	Thr 270	Ser	Ser
55				260					200					_,,		

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	Glu	Thr	Lys 275	Met	Arg	GIu	Tyr	1yr 280	Asn	гÀг	HIS	гÀг	285	GIN	Pne	Se:
	Ile	Pro		Glu	Ile	Glu	Thr		Arg	Tyr	Thr	Ser		Asn	Gln	Gl
		290					295					300				
5	Asp	Leu	Glu	Arg	Ala	Met	Ala	Asp	Pro	Asn		Glu	Val	Pro	Gly	
	305		_			310					315	_	_	_		32
	Ser	Lys	Ala	Asn	G1u 325	Lys	Iïe	GIU	Met	330	Thr	Leu	Asn	Pro	335	ΤŢ
	Ala	Gln	Val	Phe	Ile	Ser	His	Glu	Gln	Gly	Ser	Phe	Thr	Pro	Val	Me
10				340					345					350		
	Asn	Gly	Gly 355	Gly	Gly	Gln	Phe	Ile 360	Thr	Phe	Tyr	Ile	Lys 365	Glu	Lys	Arg
	Gly	Lys 370	Asn	Glu	Val	Ser	Phe 375	Ser	Gln	Ala	Lys	Gln 380	Phe	Ile	Ala	Gl
15	Lys		Val	Glu	Glu	Ser		Asp	Lys	Ile	Leu		Glu	His	Phe	Glı
	385				-	390			-		395		~			40
	Lys	Leu	Arg	Val	Lys 405	Ser	Arg	Ile	Val	Met 410	Ile	Arg	Glu			
					- • •					 ,-						
20	(2)	INF	ORMA'	rion	FOR	SEQ	ID I	NO:1	15:							
		(1)) SE	TEN	יב כז	HARA	מאינים.	rstro	7S :							
		(-				H: 1				is						
			()	B) T	PE:	amiı	no a	cid								
25			(1	D) T	OPOL	CY:	line	ear								
		(iii) MOI	LECUI	LE T	YPE:	prot	tein								
							_									
20		(iii)) HY	POTHI	ETIC	AL: :	YES									
30		(ari l	, Ob.	TCTN	AT. SC	OURCI	FT •									
		(• 1				ISM:		icoba	acte	r py:	lori					
25		(ix)) FE					_								
35						KEY:		_	ature	2						
			(1	אבו. נכ	JCAI.	ION :	.	100								
		(xi)	SE	QUEN	E DI	ESCR:	IPTIC	ON: 5	SEQ :	ID NO	0:11	5:				
40	Met	Ile	Lys	Arg	Ile	Ala	Cys	Ile	Leu	Ser	Leu	Ser	Ala	Ser	Leu	Ala
	1		•		5		-			10					15	
	Leu	Ala	Gly	Glu	Val	Asn	Gly	Phe	Phe	Met	Gly	Ala	Gly	Tyr	Gln	Glı
				20					25					30	•	
15	Gly	Arg		Gly	Pro	Tyr	Asn		Asn	Tyr	Ser	Asp		Arg	His	GL
45	•	•	35	m	a 1	*		40	*	T	<i>α</i> 1	Dh.	45	C1	Dho	λ1 :
	Asn	Asp 50	Leu	Tyr	GIY	Leu	55	Pne	гуѕ	Leu	GIY	60	Val	GIY	FILE	AIC
	Asn		Trp	Phe	Glv	Ala		Val	Tvr	Glv	Phe		Asp	Trp	Phe	Ası
	65	_,_			4-1	70	3		-1-	1	75					80
50		Ser	Gly	Thr	Glu	His	Thr	Lys	Thr	Asn	Leu	Leu	Thr	Tyr	Gly	Gly
					85					90					95	
	Gly	Gly	Asp		Ile	Val	Asn	Leu		Pro	Leu	Asp	Lys		Ala	Let
		_		100	a 3		~ 7	• .	105	a 3		~ 1	m	110	Db	D~-
55	Gly	Leu		GIY	GIY	Val	GIN		АТА	GŢĀ	Asn	Thr	Trp 125	Mec	rne	PTC
55			115					120					123			

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•	Tyr A	sp V 30	al Ası	ı Gln	Thr	Arg 135	Phe	Gln	Phe	Leu	Trp 140	Asn	Leu	Gly	Gly
	Arg M	et A	rg Val	Gly	Asp 150	Arg	Ser	Ala	Phe	Glu 155	Ala	Gly	Val	Lys	Phe 160
5	Pro M	et V	al Ası	ı Gln 165	Gly	Ser	Lys	Asp	Val 170	Gly	Leu	Ile	Arg	Tyr 175	Tyr
	Ser T	rp T	yr Va: 180	L Asp	Tyr	Val	Phe	Thr 185						1,3	
10	(2) I	NFOR	MATIO	1 FOR	SEQ	ID 1	NO:13	L6 :							
		(i)		NCE CI LENGTI TYPE:	H: 24	12 ar	nino		ls						
15				COPOL											
	(ii)	MOLEC	JLE T	YPE:	prot	tein								
20	(i	ii)	HYPOT	HETIC	AL: 1	YES									
	(vi)	ORIGII (A)	NAL SO			icoba	actei	r py	lori					
	(ix)	FEATU				_								
25				NAME/: LOCAT			_	ature	2						
	(xi)	SEQUE	NCE D	ESCR:	IPTI	: ИС	SEQ I	ID NO	0:116	5:				
30	Met L 1	ys L	ys Ph	e Phe 5	Ser	Gln	Ser	Leu	Leu 10	Ala	Leu	Ile	Ile	Ser 15	Met
	Asn A	la V	al Se	_	Met	Asp	Gly	Asn 25	Gly	Val	Phe	Leu	Gly 30	Ala	Gly
35	Tyr L		ln Gl	y Gln	Ala	Gln	Met 40	His	Ala	Asp	Ile	Asn 45	Ser	Gln	Lys
	Gln A 5	0				55					60				
	65		he Ph		70					75					80
40	Asp T			85					90					95	
			la Ala 10	0				105					110		
45	Asn A	1	15				120					125			
	1	30	yr Gl			135					140				
	Gly I 145				150					155					160
50	Ser T			165					170					175	
	Ala L		18	D				185					190		
55	Gly A		rg Le [.] 95	u Arg	Ile	Leu	Lys 200	His	Ser	Ser	Ile	Glu 205	Ala	Gly	Val

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Lys Phe Pro Met Leu Lys Lys Asn Pro Tyr Ile Thr Ala Lys Asn Leu
210 225 220

Asp Ile Gly Phe Arg Arg Val Tyr Ser Trp Tyr Val Asn Tyr Val Phe
225 230 235 240

5 Thr Phe

(2) INFORMATION FOR SEQ ID NO:117:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 256 amino acids

(B) TYPE: amino acid

(D) TOPOLOGY: linear

15 (ii) MOLECULE TYPE: protein

(iii) HYPOTHETICAL: YES

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Helicobacter pylori

(ix) FEATURE:

(A) NAME/KEY: misc feature

(B) LOCATION 1...256

25

55

20

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:117:

30 Cys Leu Phe Ser Thr Leu Gly Ala Glu His Leu Glu Gln Lys Gly Asn Tyr Ile Tyr Lys Gly Glu Glu Ala Tyr Asn Asn Lys Glu Tyr Glu Arg 40 Ala Ala Ser Phe Tyr Lys Ser Ala Ile Lys Asn Gly Glu Ser Leu Ala 35 Tyr Ile Leu Leu Gly Ile Met Tyr Glu Asn Gly Arg Gly Val Pro Lys 70 Asp Tyr Lys Lys Ala Val Glu Tyr Phe Gln Lys Ala Val Asp Asn Asp 90 40 Ile Pro Arg Gly Tyr Asn Asn Leu Gly Val Met Tyr Lys Glu Gly Lys 100 105 Gly Val Pro Lys Asp Glu Lys Lys Ala Val Glu Tyr Phe Arg Ile Ala 120 Thr Glu Lys Gly Tyr Thr Asn Ala Tyr Ile Asn Leu Gly Ile Met Tyr 45 135 Met Glu Gly Arg Gly Val Pro Ser Asn Tyr Ala Lys Ala Thr Glu Cys 150 155 Phe Arg Lys Ala Met His Lys Gly Asn Val Glu Ala Tyr Ile Leu Leu 170 50 Gly Asp Ile Tyr Tyr Ser Gly Asn Asp Gln Leu Gly Ile Glu Pro Asp 185 Lys Asp Lys Ala Val Val Tyr Tyr Lys Met Ala Ala Asp Val Ser Ser 195 200 205

Ser Arg Ala Tyr Glu Gly Leu Ser Glu Ser Tyr Arg Tyr Gly Leu Gly

215

Met Gly Tyr Ala Ser Lys Leu Ala Leu Lys Ile Cys Leu Val Gly Leu

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	Val 0					230					235					240
	Asp F	he	Asp	Ile	Asp 245	Lys	Asn	Cys	Lys	Lys 250	Lys	Asn	Thr	Ser	Ser 255	Arg
5	(2) 1	INFO	RMAT	'ION	FOR	SEQ	ID N	10:11	.8 :							
		(i)						STIC								
10			(B) TY	PE:	: 65 amin GY:	o ac		acid	ls						
		(ii)	MOL	ECUI	E TY	PE:	prot	ein								
15	(:	iii)	HYF	OTHE	TICA	L: Y	ES							,		
		(vi)				URCE SM:		i.coba	cter	py]	lori					
20		(ix)	(P		WE/K	EY:		c_fea 557	ture	:						
		(xi)	SEÇ	UENC	E DE	SCRI	PTI	ON: 5	SEQ I	D NO	0:118	3:				
25	Met i	Δrσ	LVS	Len	Phe	Ile	Pro	Leu	Leu	Leu	Phe	Ser	Ala	Leu	Glu	Ala
•	1				5					10					15	
	Asn (20					25					30		
30	Leu	Glu	Gly 35	Thr	Gln	Thr	Gln	Glu 40	Lys	Arg	His	Thr	Thr 45	Thr	ГÀЗ	Asn
	Thr '	50	Ala				55	Leu				60				
35	Ala 2	Ala	Asn	Leu	Phe	Thr 70	Asn	Ala	Glu	Ala	Ile 75	Ser	Lys	Leu	Lys	Phe 80
33	Ser	Ser	Leu	Ser			Arg	Val	Leu	Tyr 90		Tyr	Asn	Gly	Gln 95	Leu
	Thr	Ile	Glu	Asn 100	85 Phe	Leu	Pro	Tyr	Asn 105		Asn	Asn	Val	Lys 110	-	Ser
40	Phe	Thr	Asp	Ala	Gln	Gly	Asn	Val 120	Ile	Asp	Leu	Gly	Val 125	Ile	Glu	Thr
	Ile	Pro	Lys	His	Ser	Lys		Val	Leu	Pro	Gly		Ala	Phe	Asp	Ser
	Leu	130 Lys	Ile	Asp	Pro	Tyr	135 Thr		Phe	Leu	Pro	140 Lys	Ile	Glu	Ala	Thr
45	145					150					155					160
					165					170					175	
	Asn	Lys	Ile	Lys 180	Thr	Asn	Leu	Val	Val 185	Asn	Tyr	Arg	Asn	Glu 190	Asn	Lys
50	Phe	Lys	Asp 195	His	Glu	Asn	His	Trp 200	Glu	Ala	Phe	Thr	Pro 205	Gln		Ala
		210	Phe				215	Leu	Asn			220				Ser
55	Gln 225	Ser	Trp	Gly	Asp	Ala 230		Leu	Asn		Pro 235		Glu	Phe	Thr	Asn 240

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	Ser	Pro	Thr	Asp	Cys 245	Asp	Asn	Asp	Pro	Ser 250	Lys	Cys	Val	Asn	Pro 255	Gly
	Thr	Asn	Gly	Leu 260	Val	Asn	Ser	Lys	Val 265	Asp	Gln	Lys	Tyr	Val 270	Leu	Asn
5	Lys	Gln	Asp 275	Ile	Val	Asn	Lys	Phe 280	Lys	Asn	Lys	Ala	Asp 285	Leu	Asp	Val
	Ile	Val 290	Leu	Lys	Asp	Ser	Gly 295	Val	Val	Gly	Leu	Gly 300	Ser	Asp	Ile	Thr
	Pro		Asn	Asn	Asp	Asp	Gly	Lys	His	Tyr		Gln	Leu	Gly	Val	
10	305					310					315					320
	Ala	Ser	Ala	Leu	Asp 325	Pro	Lys	Lys	Leu	Phe 330	Gly	Asp	Asn	Leu	Lys 335	Thr
	Ile	Asn	Leu	Glu	Asp	Leu	Arq	Thr	Ile	Leu	His	Glu	Phe	Ser	His	Thr
				340	-		_		345					350		
15	Tare	Clv	ጥህም		Hie	Asn	Glv	Aen		Thr	Tyr	Gln	Δrα		Pro	Val
. 13	_ БУЗ	Gry_		GLY	1113		<u> </u>		1100		<u> </u>	<u> </u>		,,,,		
			355					360		_	_		365	_	_	_
	Thr	Lys 370	Asp	Gly	Gln	Val	Glu 375	Lys	Asp	Ser	Asn	380	Lys	Pro	Lys	Asp
	Ser	Asp	Gly	Leu	Pro	Tyr	Asn	Val	Cys	Ser	Leu	Tyr	Gly	Gly	Ser	Asn
20	385					390					395					400
	Gln	Pro	Ala	Phe	Pro	Ser	Asn	Tyr	Pro	Asn	Ser	Ile	Tyr	His	Asn	Cys
					405			-		410			•		415	-
		7	77 7	D		Gly	Dho	T 011	~1		TT la sa	71-	77-	17-1		Gla
	Ala	Asp	vai	420	AIA	GIY	Pne	геп	425	vai	IIII	Ala	Ата	430	пр	GIII
25	Gln	T.611	Tle		Gln	Asn	Δla	T.eu		Tle	Δen	ጥህን	ΔΊа	Asn	Leu	Glv
23	Gin	neu	435	7011	0111	7.011		440	110		11011	-7-	445			1
	Car	Gln		Acn	Tur	Asn	T.011		Δ] =	Ser	T.011	Aen		Gln	Asn	Leu
	Ser		1111	ADII	-1-	-1011		71411	71.14		ac u				P	
	_	450	_		_	_	455			_		460				G
	Ala	Asn	Ser	Met	Leu	Ser	Thr	Ile	GIn	Lys		Pne	vaı	Thr	ser	
30	465					470					475					480
	Val	Thr	Asn	His	His	Phe	Ser	Asn	Ala	Ser	Gln	Ser	Phe	Arg	Ser	Pro
					485					490					495	
	Tle	Len	Glv	٧al	Asn	Ala	Lvs	Tle	Glv	Tvr	Gln	Asn	Tvr	Phe	Asn	Asp
	114		01	500			, -		505	-] -			- 1 -	510		•
25	5 1	- 1 -	a 1		77-			a 1		T1 -	T		N		27-	Tuc
35	Pne	TIE		Leu	Ala	Tyr	TYE		116	TIE	Lys	lyr		TYL	AIG	цуз
			515					520					525		_	_
	Ala	Val	Asn	Gln	Lys	Val	Gln	Gln	Leu	Ser	Tyr	Gly	Gly	Gly	Ile	Asp
		530					535					540				
	Leu	Leu	Leu	Asp	Phe	Ile	Thr	Thr	Tyr	Ser	Asn	Lys	Asn	Ser	Pro	Thr
40	545			-		550			•		555	•				560
		Tla	Gla	Thr	Tare	Arg	Aen	Dhe	Sar	Sar		Dhe	Glv	Tle	Phe	
	GLY	116	GIII	1111		AT 9	ASII	FILE	Ser		261	FIIC	Gry			- -7
		_	_		565	_	_	_	_	570		_	_	_	575	-
	Gly	Leu	Arg	Gly	Leu	Tyr	Asn	Ser	Tyr	Tyr	Val	Leu	Asn	Lys	vai	гàг
				580					585					590		
45	Gly	Ser	Gly	Asn	Leu	Asp	·Val	Ala	Thr	Gly	Leu	Asn	Tyr	Arg	Tyr	Lys
	-		595			-		600		•			605	_	_	
	TT : -	C		The same	602	Val	Glar.		C .~	T10	Dwa	t ou		Gln	λνα	Lve
	nis		Lys	IYL	Ser	Val		116	Ser	TIE	PIO		TTE	GIII	ALG	בענג
		610					615					620				
	Ala	Ser	Val	Val	Ser	Ser	Gly	Gly	Asp	Tyr	Thr	Asn	Ser	Phe	Val	Phe
50	625					630					635					640
	Asn	Glu	Gly	Ala	Ser	His	Phe	Lys	Val	Phe	Phe	Asn	Tyr	Gly	Trp	Val
			•		645			-		650	-		-	-	655	
	Dha															
	Phe															

```
(2) INFORMATION FOR SEQ ID NO:119:
          (i) SEQUENCE CHARACTERISTICS:
               (A) LENGTH: 167 amino acids
               (B) TYPE: amino acid
5
               (D) TOPOLOGY: linear
         (ii) MOLECULE TYPE: protein
        (iii) HYPOTHETICAL: YES
10
         (vi) ORIGINAL SOURCE:
               (A) ORGANISM: Helicobacter pylori
        (ix) FEATURE:
15
               (A) NAME/KEY: misc feature
               (B) LOCATION 1...167
         (xi) SEQUENCE DESCRIPTION: SEQ ID NO:119:
20
    Met Lys Leu Val Ser Leu Ile Val Ala Leu Val Phe Cys Cys Phe Leu
                                         10
    Gly Ala Val Glu Leu Pro Gly Val Tyr Gln Thr Gln Glu Phe Leu Tyr
                                     25
    Met Lys Ser Ser Phe Val Glu Phe Phe Glu His Asn Gly Lys Phe Tyr
25
                                 40
    Ala Tyr Gly Ile Ser Asp Val Asp Gly Ser Lys Ala Lys Lys Asp Lys
                             55
    Leu Asn Pro Asn Pro Lys Leu Arg Asn Arg Ser Asp Lys Gly Val Val
30
                         70
     Phe Leu Ser Asp Leu Ile Lys Val Gly Glu Gln Ser Tyr Lys Gly Gly
                                         90
     Lys Ala Tyr Asn Phe Tyr Asp Gly Lys Thr Tyr His Val Arg Val Thr
                                     105
     Gln Asn Ser Asn Gly Asp Leu Glu Phe Thr Ser Ser Tyr Asp Lys Trp
35
     Gly Tyr Val Gly Lys Thr Phe Thr Trp Lys Arg Leu Ser Asp Glu Glu
                             135
     Ile Lys Asn Leu Lys Leu Lys Arg Phe Asn Leu Asp Glu Val Leu Lys
                                             155
40
                        150
     Thr Leu Lys Asp Ser Pro Ile
                     165
     (2) INFORMATION FOR SEQ ID NO:120:
45
          (i) SEQUENCE CHARACTERISTICS:
               (A) LENGTH: 294 amino acids
               (B) TYPE: amino acid
               (D) TOPOLOGY: linear
50
         (ii) MOLECULE TYPE: protein
        (iii) HYPOTHETICAL: YES
        (vi) ORIGINAL SOURCE:
55
```

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(A) ORGANISM: Helicobacter pylori

(ix) FEATURE:

- (A) NAME/KEY: misc_feature
- 5 (B) LOCATION 1...294

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:120:

Met Ser Asn Gln Ala Ser His Leu Asp Asn Phe Met Asn Ala Lys Asn 10 Pro Lys Ser Phe Phe Asp Asn Lys Gly Asn Thr Lys Phe Ile Ala Ile Thr Ser Gly Lys Gly Gly Val Gly Lys Ser Asn Ile Ser Ala Asn Leu Ala Tyr Ser Leu Tyr Lys Lys Gly Tyr Lys Val Gly Val Phe Asp Ala 15 55 Asp Ile Gly Leu Ala Asn Leu Asp Val Ile Phe Gly Val Lys Thr His 70 Lys Asn Ile Leu His Ala Leu Lys Gly Glu Ala Lys Leu Gln Glu Ile 20 85 90 Ile Cys Glu Ile Glu Pro Gly Leu Cys Leu Ile Pro Gly Asp Ser Gly 105 Glu Glu Ile Leu Lys Tyr Ile Ser Gly Ala Glu Ala Leu Asp Arg Phe 120 25 Val Asp Glu Glu Gly Val Leu Ser Ser Leu Asp Tyr Ile Val Ile Asp 135 140 Thr Gly Ala Gly Ile Gly Ala Thr Thr Gln Ala Phe Leu Asn Ala Ser 150 155 Asp Cys Val Val Ile Val Thr Thr Pro Asp Pro Ser Ala Ile Thr Asp 30 165 170 Ala Tyr Ala Cys Ile Lys Ile Asn Ser Lys Asn Lys Asp Glu Leu Phe 185 190 Leu Ile Ala Asn Met Val Ala Gln Pro Lys Glu Gly Arg Ala Thr Tyr 200 35 Glu Arg Leu Phe Lys Val Ala Lys Asn Asn Ile Ala Ser Leu Glu Leu 215 220 His Tyr Leu Gly Ala Ile Glu Asn Ser Ser Leu Leu Lys Arg Tyr Val 230 235 Arg Glu Arg Lys Ile Leu Arg Lys Ile Ala Pro Asn Asp Leu Phe Ser 40 245 250 Gln Ser Ile Asp Gln Ile Ala Ser Leu Leu Val Ser Lys Leu Glu Thr 265 270 Gly Thr Leu Glu Ile Pro Lys Glu Gly Leu Lys Ser Phe Phe Lys Arg

280

(2) INFORMATION FOR SEQ ID NO:121:

50 (i) SEQUENCE CHARACTERISTICS:

Leu Leu Lys Tyr Leu Gly .

290

- (A) LENGTH: 372 amino acids
- (B) TYPE: amino acid
- (D) TOPOLOGY: linear
- 55 (ii) MOLECULE TYPE: protein

	((iii)	HYI	POTHE	TIC	AL: Y	ÆS									
5		(vi)		GINA A) OF				coba	cter	py:	lori					
		(ix)	(2	ATURE A) NA B) LO	ME/F			_	ture	•						
10		(xi)	•						SEQ 1	ED NO	0:12	L:				
	Leu 1	Glu	Pro	Ser	Arg 5	Asn	Arg	Leu		His 10	Ala	Ala	Phe	Phe	Val 15	Gly
15	Leu	Phe	Ile	Val 20	Leu	Phe	Leu	Ile	Ile 25	Met	Lys	His	Gln	Thr 30	Ser	Pro
	-		35					40			Thr		45			
20		50					55				Asp	60				
	65					70					Asn 75					80
					85					90	Val				95	
25				100					105		Glu			110		
			115					120			Glu		125			
30	_	130					135				Lys	140				
	145	_				150					Ser 155					160
25					165					170	Phe				175	
35				180					185		Val Leu			190		
			195					200			Gln		205			
40		210					215				Gln	220				
	225					230					235 Phe					240
45					245					250					255	
70	_			260					265		Asp			270		
	_		275					280			His		285			
50		290					295				Lys	300				
	305					310					315 Ala					320
					-				_	-						

330

55 Pro Ser Tyr Ser Leu Asn Pro His Phe Ile Asp Ile Val Tyr Thr Tyr

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	Pro	Ala	Ile 675	Ala	Gly	Leu	Phe	Ser 680	Lys	Glu	Arg	Lys	Glu 685	Lys	Pro	Ser
		690				Asp	695	_				700		-		_
5	Tyr 705	Glu	Lys	Ala	His	Lys 710	Ile	Ile	Pro	Ile	Ser 715	Thr	Arg	Ile	His	Ala 720
	Lys	Asp	Val	Val	Leu 725	Ile	Tyr	Lys	Lys	Met 730	Pro	Phe	Pro	Leu	Glu 735	Asn
10	Leu	Asp	Ile	Val 740	Ala	Gln	Asp	Asp	Arg 745	Val	Lys	Ile	Asp	Gly 750	Asn	Tyr
	Lys	Asn	Ala 755	Met	Ile	Met	Ala	Asp 760	Leu	۷al	His	Gly	Ala 765	Leu	Tyr	Leu
	-	770				Ser	775	_	-			780				_
15	Asp	Phe	Val	Glu	Gly	Gly	Leu	Phe	Thr	Leu	Ile	Gly	Ala	Leu	Glu	Asp
	785					790			•		795					800
	Gln	Val	Phe	Asn	Gly 805	Glu	Leu	Lys	Phe	Gln 810	Asn	Thr	Ser	Leu	Lys 815	Asn
20	Phe	Ala	Leu	Met 820	Gln	Asn	Met	Val	Asn 825	Leu	Ile	Asn	Thr	Ile 830	Pro	Ser
	Leu	Ile	Val 835	Phe	Arg	Asn	Pro	His 840	Leu	Gly	Ala	Asn-	Gly 845	Tyr	Gln	Ile
	Lys	Thr 850	Gly	Ser	Val	Val	Phe 855	Gly	Ile	Thr	Lys	Glu 860	Tyr	Leu	Gly	Leu
25	Glu 865	Lys	Ile	qaA	Leu	Val 870	Gly	Lys	Thr	Leu	Asp 875	Ile	Ala	Gly	Asn	Gly 880
	Ile	Ile	Glu	Leu	Asp 885	Lys	Asn	Lys	Leu	Asp 890	Leu	Asn	Leu	Glu	Val 895	Ser
30	Thr	Ile	Lys	Ala 900	Leu	Ser	Asn	Val	Leu 905	Asn	Lys	Ile	Pro	Ile 910	Val	Gly
	Tyr	Leu	Val 915	Leu	Gly	Lys	Gly	Gly 920	Lys	Ile	Thr	Thr	Asn 925	Val	Asn	Val
	Lys	Gly 930	Thr	Leu	Asp	Lys	Pro 935	Lys	Thr	Gln	Val	Thr 940	Leu	Ala	Ser	Asp
35	Ile	Ile	Gln	Ala	Pro	Phe	Lys	Ile	Leu	Arg	Arg	Ile	Phe	Thr	Pro	Ile
	945					950					955					960
	Asp	Ile	Ile	Val	Asp 965	Glu	Val	Lys	Lys	Asn 970	Ile	Asp	Ser	Lys	Arg 975	Lys
40	Leu	Lys														

(2) INFORMATION FOR SEQ ID NO:123:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 477 amino acids

(B) TYPE: amino acid

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

(iii) HYPOTHETICAL: YES

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Helicobacter pylori

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(ix) FEATURE:

- (A) NAME/KEY: misc_feature
- (B) LOCATION 1...477

5 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:123:

	Met 1	Asn	Thr	Ile	Ile 5	Arg	Tyr	Ala	Ser	Leu 10	Trp	Gly	Leu	Cys	Ile 15	Thr
10	Leu			20					25					30	Lys	
			35					40					45		Pro	
		50					55					60			Lys	
15	65					70					75				Lys	80
	_	-			85					90					Thr 95	
20				100					105					110	Met Ala	
			115					120					125		His	
25	-	130		_			135					140			Asn	
23	145					150					155				Arg	160
					165					170					175 Gln	
30				180					185					190	Ile	
			195					200					205		Val	
35		210					215					220			Ser	
75	225					230					235				Gln	240
	_				245					250					255 Leu	
40				260					265					270	Gln	
		_	275					280					285		Ala	
45	_	290					295					300			Val	
.5	305					310	,				315					320 Arg
					325					330					335	Ala
50				340					345					350		Ser
			355					360					365		Lys	
55		370					375					380				Ile
رر	Leu	GIU	GIII	GIU	пур	vab	- Lu		u	- X -	713	د ړ ــ			<u>F</u>	

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340 345 Asn Arg Ser His Ile Lys His Ile Arg Phe Asn Met Ala Tyr Leu Asn 355 360 Ser Leu Leu Lys 5 370 (2) INFORMATION FOR SEQ ID NO:122: (i) SEQUENCE CHARACTERISTICS: 10 (A) LENGTH: 978 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (ii) MOLECULE TYPE: protein 15 (iii) HYPOTHETICAL: YES (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori 20 (ix) FEATURE: (A) NAME/KEY: misc feature (B) LOCATION 1...978 25 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:122: Met Lys Lys Arg Lys His Val Ser Lys Lys Val Phe Asn Val Ile Ile 10 Leu Phe Val Ala Val Phe Thr Leu Leu Val Val Ile His Lys Thr Leu 30 25 Ser Asn Gly Ile His Ile Gln Asn Leu Lys Ile Gly Lys Leu Gly Ile 40 Ser Glu Leu Tyr Leu Lys Leu Asn Asn Lys Leu Ser Leu Glu Val Glu 55 35 Arg Val Asp Leu Ser Ser Phe Phe His Gln Lys Pro Thr Lys Lys Arg 75 Leu Glu Val Ser Asp Leu Ile Lys Asn Ile Arg Tyr Gly Ile Trp Ala Val Ser Tyr Phe Glu Lys Leu Lys Val Lys Glu Ile Ile Leu Asp Asp 40 105 Lys Asn Lys Ala Asn Ile Phe Phe Asp Gly Asn Lys Tyr Glu Leu Glu 120 Phe Pro Gly Ile Lys Gly Glu Phe Ser Leu Glu Asp Asp Lys Asn Ile 135 140 45 Lys Leu Lys Ile Ile Asn Leu Leu Phe Lys Asp Val Lys Val Gln Val 150 155 Asp Gly Asn Ala His Tyr Ser Pro Lys Ala Arg Lys Met Ala Phe Asn 165 170 Leu Ile Val Lys Pro Leu Val Glu Pro Ser Ala Ala Ile Tyr Leu Gln 50 . 185 190 Gly Leu Thr Asp Leu Lys Thr Ile Glu Leu Lys Ile Asn Thr Ser Pro 200 Met Lys Ser Leu Ala Phe Leu Lys Pro Leu Phe Gln Arg Gln Ser Gln 215 220 55

Lys Asn Leu Lys Thr Trp Ile Phe Asp Lys Ile Gln Phe Ala Ser Phe

	225					230					235					240
	225 Lys	Ile	Asp	Asn	Ala 245	Leu	Ile	Lys	Ala	Asn 250	Phe	Thr	Pro	Ser	Glu 255	Phe
ح	Ile	Pro	Ser	Leu 260	Leu	Glu	Asn	Ser	Val 265		Lys	Ala	Thr	Leu 270	Ile	Lys
5	Pro	Ser	Val 275	Val	Phe	Asn	Asp	Gly 280		Ser	Pro	Ile	Lys 285	Met	Asp	Lys
	Thr		Leu	Ile	Phe	Lys	Asn 295		Gln	Leu	Leu	Ile 300	Gln	Pro	Gln	Lys
10		290 Thr	Tyr	Glu	Thr	Met 310		Leu	Thr	Gly	Ser 315		Ala	Thr	Phe	Ser 320
	305 Asn	Leu	Leu	Glu	Ala 325	Pro	Lys	Leu	Glu	Val 330		Leu	Lys	Thr	Thr 335	Pro
15	Asn	Tyr	Tyr	Gly 340	Asp	Ser	Ile	Lys	Asp 345	Leu	Leu	Ser	Ala	Tyr 350	Lys	Val
13	Val	Leu	Pro	Leu	Asp	Lys	Ile	Ser 360	Met	Pro	Ser	Ser	Ala 365	Asp	Leu	Lys
		370	Leu	Gln			375					380				
20	385	Ser		Asn		390					395					400
	Leu			Gln	405					410					415	
25				Tyr 420					425					430		
			435	Ala				440					445			
		450		Leu			455					460				
30	465			Ser		470					475					400
				Leu	485					490					495	
35				Glu 500					505					510		
			515	Asp				520					525			
		530		Ser			535					540				
40	545			Ser		550					555					200
					565					570					5/5	
45				Met 580			-		585					590		
			595	Asp				600)				605			
50		610					615					620				Phe
50	625					630					635					Pro 640 Thr
					645	;				650					633	
55	Leu	Asn	Asn	660		Leu	sel	. TTE	665		FIIC	, Jieu	up	670	-1-	Met

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385 390 395 Ala Arg Ala Lys Ile Glu Ser Ser Lys Ala Ser Leu Asp Ala Ala Asn 405 410 Leu Ser Phe Ala Asn Ile Lys Arg Lys Tyr Asp Ala Asn Leu Val Asp 5 425 Phe Thr Thr Tyr Leu Arg Gly Leu Thr Thr Arg Phe Asp Ala Glu Val 440 Ala Tyr Asn Leu Ala Leu Asn Asn Tyr Glu Val Gln Lys Ala Asn Tyr 10 Ile Phe Asn Ser Gly His Lys Ile Asp Asp Tyr Val His 470 (2) INFORMATION FOR SEQ ID NO:124: 15 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 412 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear 20 (ii) MOLECULE TYPE: protein (iii) HYPOTHETICAL: YES (vi) ORIGINAL SOURCE: 25 (A) ORGANISM: Helicobacter pylori (ix) FEATURE: (A) NAME/KEY: misc feature (B) LOCATION 1...412 30 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:124: Met Leu Ser Phe Ile Ser Ala Phe Asp Lys Arg Gly Val Ser Ile Arg 10 35 Leu Leu Thr Ala Leu Leu Leu Phe Ser Leu Gly Leu Ala Lys Asp Leu Glu Ile Gln Thr Phe Val Ala Lys Tyr Leu Ser Lys Asn Gln Lys Ile Gln Ala Leu Gln Glu Gln Ile Asp Ala Leu Asp Ser Gln Glu Lys 40 Val Val Ser Lys Trp Asp Asn Pro Ile Leu Tyr Leu Gly Tyr Asn Asn 70 Ala Asn Val Ser Asp Phe Phe Arg Leu Asp Ser Thr Leu Met Gln Asn 90 45 Met Ser Leu Gly Leu Ser Gln Lys Val Asp Leu Asn Gly Lys Lys Leu 105 Thr Gln Ser Lys Met Ile Asn Leu Glu Lys Gln Lys Lys Ile Leu Glu 120 Leu Lys Lys Thr Lys Gln Gln Leu Val Ile Asn Leu Met Ile Asn Gly 50 135 140 Ile Glu Asn Tyr Lys Asn Gln Glu Ile Glu Leu Leu Asn Thr Ala 155 Ile Lys Asn Leu Glu Asn Thr Leu Tyr Gln Ala Asn His Ser Ser Ser 170 55 Pro Asp Leu Ile Ala Ile Ala Lys Leu Glu Ile Leu Lys Ser Leu Leu

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				180					185					190		
			195	Lys				200			Leu		205			
5		210	_				215				Glu	220				
	Pro 225	Lys	Asn	Phe	Glu	Phe 230	Asn	Asn	Glu	Gln	Glu 235	Leu	His	Asn	Ile	Ser 240
	Ala	Thr	Asn	Tyr	Asp 245	Ile	Ala	Ile	Ala	Arg 250	Leu	Asp	Glu	Glu	Lys 255	Ala
10	Gln	Lys	Asp	Ile 260	Thr	Leu	Ala	Lys	Lys 265	Ser	Phe	Leu	Glu	Asp 270	Ile	Asn
			275					280			Gln		285			
15		290					295				Pro	300				
	Ala 305	Lys	Leu	Val	Glu	Gln 310	Lys	Lys	Lys	Glu	Ser 315	Leu	Ala	Phe	Lys	Ser 320
	Glu	Val	Glu	Asn	Ala 325	Lys	Asn	Lys	Thr	Arg 330	His	Leu	Ala	Leu	Lys 335	Leu
20	Leu	Lys	Lys	Leu 340	Glu	Thr	Leu	Gln	Lys 345	Asn	Leu	Glu	Ser	Ile 350	Asn	Lys
			355					360			Ile		365			
25	_	370					375				Asn	380				
	Ile 385	Thr	Ile	Gln	Ile	Thr 390	Gln	Leu	Glu	Thr	Leu 395	Ser	Ala	Leu	Asn	Ser 400
	Ala	Tyr	Leu	Ser	Leu 405	Gln	Asn	Leu	Lys	Gly 410	Leu	Glu				
30	(2)	INFO	ORMA!	TION	FOR	SEQ	ID I	NO:1	25:							
		(i)) SEC	QUEN	CE CI	HARA	CTER:	ISTI	CS:							
35				A) Li B) T					acio	is						
			(1	D) T	OPOL	OGY:	lin	ear								
		(ii)) MO	LECU	LE T	YPE:	pro	tein								
40		(iii)	HY:	POTH	ETIC	AL: `	YES									
		(vi)		IGINA A) O				icob	acte:	r py	lori					
45		(ix)	(ATUR: A) N: B) L	AME/				atur	e						
50		(xi)) SE	QUEN	CE D	ESCR	IP TI	ON:	SEQ	ID N	0:12	5:				
J0	Met 1	Arg	Ile	Val	Arg 5	Asn	Leu	Phe	Leu	Val 10	Ser	Phe	Val	Ala	Tyr 15	Ser
		Ala	Phe	Ala 20		Asp	Leu	Glu	Thr 25	Gly	Thr	Lys	Asn	Asp 30	Lys	Lys
55	Ser	Glv	Lvs		Phe	Tvr	Lys	Leu	His	Lys	Asn	His	Gly	Ser	Glu	Thr

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35 40 Glu Thr Lys Asn Asp Lys Lys Leu Tyr Asp Phe Thr Lys Asn Ser Gly 55 60 Leu Glu Gly Val Asp Leu Glu Lys Ser Pro Asn Leu Lys Ser His Lys 5 Lys Ser Asp Lys Lys Phe Tyr Lys Gln Leu Ala Lys Asn Asn Ile Ala 90 Glu Gly Val Ser Met Pro Ile Val Asn Phe Asn Lys Ala Leu Ser Phe 105 Gly Pro Tyr Phe Glu Arg Thr Lys Ser Lys Lys Thr Gln Tyr Met Asp 10 120 Gly Gly Leu Met Met His Ile Arg Phe 130 15 (2) INFORMATION FOR SEQ ID NO:126: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 309 amino acids (B) TYPE: amino acid 20 (D) TOPOLOGY: linear (ii) MOLECULE TYPE: protein (iii) HYPOTHETICAL: YES 25 (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori (ix) FEATURE: 30 (A) NAME/KEY: misc feature (B) LOCATION 1...309 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:126: Leu Met Pro Gln Asn Gln Leu Val Ile Thr Ile Ile Asp Glu Ser Gly Ser Lys Gln Leu Lys Phe Ser Lys Asn Leu Lys Arg Asn Leu Ile Ile Ser Val Val Ile Leu Leu Ile Val Gly Leu Gly Val Gly Phe Leu 40 Lys Phe Leu Ile Ala Lys Met Asp Thr Met Thr Ser Glu Arg Asn Ala Val Leu Arg Asp Phe Arg Gly Leu Tyr Gln Lys Asn Tyr Ala Leu Ala 45 Lys Glu Ile Lys Asn Lys Arg Glu Glu Leu Phe Ile Val Gly Gln Lys 90 Ile Arg Gly Leu Glu Ser Leu Ile Glu Ile Lys Lys Gly Ala Asn Gly 105 Gly Gly His Leu Tyr Asp Glu Val Asp Leu Glu Asn Leu Ser Leu Asn 50 120 Gln Lys His Leu Ala Leu Met Leu Ile Pro Asn Gly Met Pro Leu Lys 135 140 Thr Tyr Ser Ala Ile Lys Pro Thr Lys Glu Arg Asn His Pro Ile Lys 150 155 55 Lys Ile Lys Gly Val Glu Ser Gly Ile Asp Phe Ile Ala Pro Leu Asn

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					165					170					175	
	Thr	Pro	Val	Tyr 180	Ala	Ser	Ala	Asp	Gly 185	Ile	Val	Asp	Phe	Val 190	Lys	Thr
5	Arg	Ser	Asn 195	Ala	Gly	Tyr	Gly	Asn 200	Leu	Val	Arg	Ile	Glu 205	His	Ala	Phe
	-	210		Ser			215					220				
	225			Ile		230					235					240
10	-			Gly	245					250					255	
	-			Asp 260					265					270		
15			275	Leu				280					285			
	-	290		Glu		Ile	Val 295	GIn	Leu	GIn	Glu	300	vaı	Asp	rys	Asp
20	Thr 305	Leu	Lys	Gly	Gln											
20	(2)	INF	ORMA?	rion	FOR	SEQ	ID 1	NO:1	27:							
25			(2 (1 (1	QUENCA) LIB) TIO) TO	ENGTI YPE : OPOLO	H: 3: amin OGY:	32 am no ao lino	mino cid ear		ds						
30		(iii)) HY	ротні	ETIC	AL:	YES									
		(vi		IGINZ A) O				icob	acte:	r py	lori					
35		(ix	(2	ATURI A) Ni B) L	AME/				atur	e						
40				QUEN												
	1			Phe	5					10					15	
				Lys 20					25					30		
45	_		35	His				40					45			
		50		Ala			55					60				
50	65			Phe		70					75					80
				Asp	85					90					95	
				Val 100					105					110		
55	Val	Ser	Asp	Phe	Ser	Pro	Leu	Phe	Ser	Leu	Pro	Tyr	Phe	Ile	Ala	Phe

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			115					120					125			
	Leu	Phe 130	Ala	Ile	Phe	Met	Leu 135	Val	Gly	Ile	Ser	Asn 140	Ala	Ile	Asn	Ile
5			Gly	Phe	Asn	Gly 150	Leu	Ala	Ser	Gly	Ile 155	Cys	Ala	Ile	Ala	Leu 160
3	145 Leu	Val	Ile	His		Ile	Asp	Pro	Ser			Ser	Cys	Leu		
	Tyr	Met	Val	Leu	165 Gly	Phe	Met	Val	Leu	170 Asn	Phe	Pro	Ser	Gly	175 Lys	Ile
10	Phe	Leu	Gly	180 Asp	Gly	Gly	Ala	Tyr	185 Phe	Leu	Gly	Leu	Val	190 Cys	Gly	Ile
	Sar	Leu	195	His	Leu	Ser	Leu	200 Glu	Gln	Lvs	Ile	Ser	205 Val	Phe	Phe	Glv
		210					215			_		220				
15	225					Tyr 230					235					240
	Arg	Arg	Lys	Ile	Lys 245	Arg	Gln	Lys	Ala	Thr 250	Met	Pro	Asp	Asn	Leu 255	His
	Leu	His	Thr	Leu 260	Leu	Phe	Lys	Phe	Leu 265	Gln	Gln	Arg	Ser	Phe 270	Asn	Tyr
20	Pro	Asn	Pro 275	Leu	Cys	Ala	Phe	Ile 280	Leu	Ile	Leu	Cys	Asn 285	Leu	Pro	Phe
	Ile	Leu 290		Ser	Val	Leu	Phe 295		Leu	Asp	Ala	Tyr 300		Leu	Ile	Val
25			Leu	Val	Phe			Cys	Tyr	Leu			Tyr	Ala	Tyr	Leu
25	305 Asn	Arg	Gln	Val	Cys	310 Ala	Leu	Glu	Lys	Arg	315 Ala	Phe				320
					325					330						
30	(2)	INF	ORMA:	rion	FOR	SEQ	ID 1	NO:12	28:							
50		(i)				IARAC				ic						
			(1	3) T	PE:	amir	o ac	cid	acit							
35			(I) T(OPOLO	OGY:	line	ear								
		(ii)	MOI	LECUI	LE TY	PE:	prot	cein		•						
	((iii)	HYI	POTH	ETIC	AL: Y	ŒS									
40		(vi)				URCE		b .		1						
						(SM:	neri	LCOD	ccei	. py	LOII					
		(ix)		ATURI A) NA		ŒY:	misc	_fea	ture	<u> </u>						
45			(I	3) LO	CAT	ON 1	L2	271								
		(xi)	SEC	QUENC	CE DE	ESCRI	PTIC	ON: S	SEQ]	D NC	128	3:				
50		Asn	Ile	Phe		Arg	Ile	Ile	Cys		Thr	Ala	Ile	Val	Leu 15	Gly
50	1 Phe	Phe	Asn		5 Leu	Asp	Ala	Lys		10 His	Lys	Glu	Lys			Asp
	His	Lys	Ile	20 Thr	Arg	Glu	Leu	Lys	25 Val	Gly	Ala	Asn	Pro	30 Val	Pro	His

35 40 45 55 Ala Gln Ile Leu Gln Ser Val Val Asp Asp Leu Lys Glu Lys Gly Ile

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		50					55					60			•	
	Lys 65	Leu	Val	Ile	Val	Ser 70	Phe	Thr	Asp	Tyr	Val 75	Leu	Pro	Asn	Leu	Ala 80
5	Leu	Asn	Asp	Gly	Ser 85	Leu	Asp	Ala	Asn	Tyr 90	Phe	Gln	His	Arg	Pro 95	Tyr
,	Leu	Asp	Arg	Phe 100	Asn	Leu	Asp	Arg	Lys 105	Met	His	Leu	Val	Gly 110	Leu	Ala
	Asn	Ile	His 115		Glu	Pro	Leu	Arg 120	Phe	Tyr	Ser	Gln	Lys 125	Ile	Thr	Asp
10	Ile	Lys 130		Leu	Lys	Lys	Gly 135	Ser	Val	Ile	Ala	Val 140	Pro	Asn	Asp	Pro
	Ala 145	Asn	Gln	Gly	Arg	Ala 150	Leu	Ile	Leu	Leu	His 155	Lys	Gln	Gly	Leu	Ile 160
15	Ala	Leu			165					170					175	
	_	Asn		180					185					190		
		Lys	195					200					205			
20		Leu 210					215					220				
	225	Pro				230					235					240
25		Ala			245					250					255	Arg
	Lys	Phe	Ile	Leu 260	Asp	Thr	Tyr	Lys	Gly 265	Ala	Ile	Ile	Pro	Ala 270	Phe	
	(2)	INFO	ORMA!	rion	FOR	SEQ	ID !	NO:1	29:							
30		(i)		QUENC						de						
			(1		YPE:	ami	no a	cid	ac.i.	45						
35		(33)		LECUI												
		(iii)														
40				IGIN												
,,		,		A) O				icob	acte	r py	lori					
		(ix		ATUR: A) N.		KEY:	mis	c_fe	atur	e				٠		
45				B) L												
				QUEN												
50	1				5					10					15	Leu
				20					25					30		Ala
	Thr	Ser	Asn	Pro	Ser	Leu	Phe	Cys	Glu	Ala	Ile	Thr	Lys	Ser	Ala	Phe

55 Tyr Gln Asp Glu Ile Ala Lys Leu Lys Gly Lys Lys Ala Lys Glu Ile

		50	Thr	T 011	ת דת	T 011	55	λαπ	Tla	Lan	C1n	60	5ax	Sar	772	T.011
	65					70					75					80
5	Met	Pro	Leu	Tyr	Glu 85	Lys	Asp	Pro	Asn	Asn 90	Gly	Tyr	Ile	Ser	Leu 95	Glu
	Ile	Asp	Pro	Phe 100	Leu	Glu	Asp	Asp	Ala 105	Ile	Lys	Ser	Ile	Asp 110	Glu	Ala
	Lys	Arg	Leu 115	Phe	Lys	Thr	Leu	Asn 120	Arg	Pro	Asn	Val	Met 125	Ile	Lys	Val
10	Pro	Ala 130	Ser	Glu	Ser	Ala	Phe 135		Val	Ile	Ser	Ala 140		Ala	Gln	Ala
	Ser		Pro	Ile	Asn	Val		Leu	Val	Phe	Ser		Lys	Ile	Ala	Gly
	145					150					155				_	160
15			Ala		165					170					175	
	Ser	⁻Va`l⁻	-Phe	Val 180	Ser	Arg	Phe	Asp	Lys 185	Glu	Ile	Asp	Pro	Leu 190	Vāl	Pro
	Gln	Asn	Leu 195	Gln	Ala	Gln	Ser	Gly 200	Ile	Met	Asn	Ala	Thr 205	Glu	Cys	Tyr
20	Tyr	Gln 210	Ile	Asn	Gln	His	Ala 215	Asn	Lys	Leu	Ile	Ser 220	Thr	Leu	Phe	Ala
	Ser	Thr	Gly	Val	Lys	Ser	Asn	Ser	Leu	Ala	Lys	Asp	Tyr	Tyr	Ile	Lys
	225					230	_				235	_	_	_		240
25	Ala	Leu	Cys	Phe	Lys 245	Asn	Ser	Ile	Asn	Thr 250	Ala	Pro	Leu	Asp	A1a 255	Leu
			Tyr	260					265		_			270		
	Ile	Thr	Glu 275	Ile	Glu	Ala	Phe	Lys 280	Lys	Glu	Leu	Lys	Thr 285	His	Asn	Ile
30	Asp	Leu 290	Glu	Asn	Thr	Ala	Gln 295	Lys	Leu	Leu	Lys	Glu 300	Gly	Leu	Ile	Ala
	Phe 305	Lys	Gln	Ser	Phe	Glu 310	Lys	Leu	Leu	Ser	Ser 315	Phe				
35	(2)	TNFC	RMAT	NOI	FOR	SEO	ID N	JO:13	30:							
	(-/		SEC													
		(1)	_	_					ació	is						
40						amir OGY:										
		(ii)	MOI	LECUI	E TY	PE:	prot	ein								
	((iii)	нуі	POTHE	TICA	AL: Y	ES									
45		(vi)	ORI					.coba	acter	py]	.ori					
50		(ix)		A) NA	ME/K	EY:		_	iture	:						
		(xi)	SEÇ	UENC	E DE	SCRI	PTIC	N: S	EQ I	D NC	:130):				

Met Lys Thr Asn Gly His Phe Lys Asp Phe Ala Trp Lys Lys Cys Phe

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	1				5					10					15	
		_		20					25		Gly			30		
5			35					40			Asn		45			
		50					55				Leu	60		•		
	Phe 65	Gln	Tyr	Ser	Asp	Asn 70	Ile	Ala	Lys	Glu	Tyr 75	Glu	Asn	Lys	Phe	Lys 80
10					85					90	Leu				95	
	-			100					105		Asp			110		
15		_	115					120			Met		125			
		130					135				Lys	140	•			
	Leu 145	Leu	Phe	Ser	Thr	Gly 150	Leu	Asp	Lys	Met	Glu 155	Arg	Val	Leu	Ile	Pro 160
20		Gly	Phe	Val	Lys 165	Val	Thr	Ile	Leu	Glu 170	Pro	Met	Ser	Gly	Glu 175	Ser
	Leu	Asp	Ser	Phe 180	Thr	Met	Asp	Leu	Ser 185	Glu	Leu	Asp	Ile	Gln 190	Glu	Lys
25	Phe	Leu	Lys 195	Thr	Thr	His	Ser	Ser 200	His	Ser	Gly	Gly	Leu 205	Val	Ser	Thr
	Met	Val 210	Lys	Gly	Thr	Asp	Asn 215	Ser	Asn	Asp	Ala	Ile 220	Lys	Ser	Ala	Leu
	Asn 225		Ile	Phe	Ala	Ser 230	Ile	Met	Gln	Glu	Met 235	Asp	Lys	Lys	Leu	Thr 240
30		Arg	Asn	Leu	Glu 245		Tyr	Gln	Lys	Asp 250	Ala	Lys	Glu	Leu	Lys 255	Asn
	Lys	Arg	Asn	Arg 260												
35	(2)	INFO	ORMA!	rion	FOR	SEQ	ID I	NO:1	31:							
		(i)				HARAG				ids						
40				•		ami: OGY:										
		(ii)	MOI	LECU	LE T	YPE:	pro	tein								
45		(iii)	HY	POTH	ETIC	AL: `	YES									
45		(vi)				OURC		icob	acte:	r py:	lori					
50		(ix)		A) N	AME/	KEY:		_	atur	e						
		(xi) SE	QUEN	CE D	ESCR:	IPTI	ON:	SEQ	ID N	0:13	1:				
55	Leu	Asn	Phe	Asn	Asn	Leu	Thr	Ala	Asn	Gly	Ala	Leu	Asn	Phe	Asn	Gly

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	Tyr	Ala	Pro	Ser 20	Leu	Thr	Lys	Ala	Leu 25	Met	Asn	Val	Ser	Gly 30	Gln	Phe
5	Val	Leu	Gly 35	Asn	Asn	Gly	Asp	Ile 40	Asn	Leu	Ser	Asp	Ile 45	Asn	Ile	Phe
	Asp	Asn 50	Ile	Thr	Lys	Ser	Val 55	Thr	Tyr	Asn	Ile	Leu 60	Asn	Ala	Gln	Lys
	Gly 65	Ile	Thr	Gly	Ile	Ser 70	Gly	Ala	Asn	Gly	Tyr 75	Glu	Lys	Ile	Leu	Phe 80
10	Tyr	Gly	Met	Lys	Ile 85	Gln	Asn	Ala	Thr	Tyr 90	Ser	Asp	Asn		Asn 95	Ile
	Gln	Thr	Trp	Ser 100	Phe	Ile	Asn	Pro	Leu 105	Asn	Ser	Ser	Gln	Ile 110	Ile	Gln
15	Glu	Ser	Ile 115	Lys	Asn	Gly	Asp	Leu 120	Thr	Ile	Glu	Val	Leu 125	Asn	Asn	Pro
	Asn	Ser 130	Ala	Ser	Asn	Thr	Ile 135	Phe	Asn	Ile	Ala	Pro 140	Glu	Leu	Tyr	Asn
	Tyr 145	Gln	Asp	Ser	Lys	Gln 150	Asn	Pro	Thr	Gly	Tyr 155	Ser	Tyr	Asp	Tyr	Ser 160
20	Asp	Asn	Gln	Ala	Gly 165	Thr	Tyr	Tyr	Leu	Thr 170	Ser	Asn	Ile	Lys	Gly 175	Leu
	Phe	Thr	Pro	Lys 180	Gly	Ser	Gln	Thr	Pro 185	Gln	Thr	Pro	Gly	Thr 190	Tyr	Ser
25	Pro	Phe	Asn 195	Gln	Pro	Leu	Asn	Ser 200	Leu	Asn	Ile	Tyr	Asn 205	Lys	Gly	Phe
	Ser	Ser 210	Glu	Asn	Leu	Lys	Thr 215	Leu	Leu	Gly	Ile	Leu 220	Ser	Gln	Asn	Ser
		Thr	Leu	Lys	Glu	Met	Ile	Glu	Ser	Asn		Leu	Asp	Asn	Ile	
20	225	- 1 -	3	a 1	**- 1	230	~ 1_	*	T	3	235	T1 -	T	-1 -	mb	240
30					245	Leu				250					255	
			_	260		Leu			265					270		
35			275			Asn		280					285			
		290				Ser	295					300				
	305	Ser	ser	PIO	Cys	Ala 310	reu	ASP	Ser	ALA	315	Cys	ser	ser	PHE	320
40	Asn	Thr	Tyr	Leu	Gly 325	Gln	Leu	Leu	Gly	Ser 330	Thr	Ser	Pro	Tyr	Leu 335	Gly
	_			340	_	Phe	_		345			_		350		
45			355			Ala	-	360					365			
		370				Asn	375				-	380				
	Gln 385	Ala	Thr	Asp	Asn	Ile 390	Phe	Asn	Leu	Leu	Gly 395	Gln	Glu	Gly	Ile	Asp 400
50		Ile	Phe	Asn	Gln 405	Gly	Asn	Leu	Ala	Asn 410		Leu	Ser	Gln	Met 415	
	Met	Glu	Lys	Ile 420		Gln	Ala	Gly	Gly 425		Gly	Asn	Phe	Ile 430		Asn
55	Ala	Leu	Ser 435		Leu	Ser	Lys	Glu 440		Pro	Ala	Ser	Leu 445		Asp	Glu

	Thr	Leu 450	Gly	Gln	Leu	Ile	Gly 455	Gln	Asn	Asn	Leu	Asp 460	Asp	Leu	Leu	Asr
	Asn 465	Ser	Gly	Val	Met	Asn 470	Glu	Ile	Gln	Asn	Ile 475	Ile	Ser	Gln	Lys	Let 480
5		Ile	Phe	Gly	Asn 485	Phe	Val	Thr	Pro	Ser 490		Ile	Glu	Asn	Tyr 495	Lev
	Ala	Lys	Gln	Ser 500		Lys	Ser	Met	Leu 505		Asp	Lys	Gly	Leu 510	Leu	Asn
10	Phe	Ile	Gly 515		Tyr	Ile	Asp	Ala 520	Ser	Glu	Leu	Ser	Ser 525	Ile	Leu	Gly
	Val	Ile 530	Leu	Lys	Asp	Ile	Thr 535	Asn	Pro	Pro	Thr	Ser 540	Leu	Gln	Lys	Asp
	Ile 545	Gly	Val	Val	Ala	Asn 550	Asp	Leu	Leu	Asn	Glu 555	Phe	Leu	Gly	Gln	Asp 560
15	Val	Val	Lys	Lys	Leu 565	Glu	Ser	Gln	Gly	Leu 570			Asn	Ile	Ile 575	Asn
	Asn	Val	Ile	Ser 580	Gln	Gly	Gly	Leu	Ser 585	Gly	Val	Tyr	Asn	Gln 590	Gly	Leu
20	Gly	Ser	Val 595	Leu	Pro	Pro	Ser	Leu 600	Gln	Asn	Ala	Leu	Lys 605	Glu	Asn	Asp
		610				Ser	615					620				
	Gly 625	Tyr	Phe	Asn	Phe	Leu 630	Ser	Asn	Gly	Tyr	Val 635	Phe	Val	Asn	Asn	Ser 640
25	Ser	Phe	Ser	Asn	Ala 645	Thr	Gly	Gly	Ser	Leu 650	Asn	Phe	Val	Ala	Asn 655	Lys
				660		Gly			665					670		
30	-		675			Ala		680					685			
		690				Asn	695					700				
	705					Gly 710					715					720
35			-		725	Ser				730					735	
				740		His			745					750		
40			755	_		Ile		760					765			
		770				Leu	775					780				
	785					Ala 790					795					800
45		_	-		805	Thr				810					815	
				820		Val			825					830		
50			835			His		840					845			
	_	850				Met	855					860				
	865			•		Val 870					875					880
55	Ala	Ile	Tvr	Tyr	Gly	Tyr	Asn	Asn	Gln	Ile	Thr	Gly	Gly	Ser	Ser	Leu

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					885					890					895	
	Asp	Asn	Tyr	Leu 900	Lys	Leu	Tyr	Ala	Leu 905		Asp	Ile	Asn	Gly 910		His
5	Met	Val	Met 915	Thr	Asp	Asn	Gly	Leu 920		Tyr	Asn	Gly	Gln 925	Ala		Ser
	Val	Lys 930	_	Gly	Gly	Leu	Val 935	Val	Gly	Phe	Lys	Asp 940	Ser	Gln	Asn	Gln
	Tyr 945		Tyr	Thr	Ser	Ile 950	Leu	Tyr	Asn	Lys	Val 955	Lys	Ile	Ala	Val	Ser 960
10	Asn	Asp	Pro	Ile	Asn 965	Asn	Pro	Gln	Ala	Pro 970	Thr	Leu	Lys	Gln	Tyr 975	Ile
	Ala	Gln	Ile	Gln 980	Gly	Val	Gln	Ser	Val 985	Asp	Ser	Ile	Asp	Gln 990	Ala	Gly
15	Gly	Asn	Gln 995	Ala	Ile	Asn	Trp	Leu 100		Lys	Ile	Phe	Glu 100	Thr 5	Lys	Gly
	Ser	-Pro-	Leu	-Phe	Ala	Pro	Tyr 101	Tyr		Glü	Ser	His	Ser	Thr	Lys	Asp
	T.611			Tle	ΔΊа	Glv			Δla	Δsn	Thr			Val	Tle	Δla
	102					103	_				103		014			104
20	Asn	Pro	Asn	Phe	Lys 104		Asp	Ala	Thr	Asn 105		Leu	Gln	Ile	Asn 105	
	Tyr	Thr	Gln	Gln 1060		Ser	Arg	Leu	Ala 1065	_	Leu	Ser	Asp	Thr 107		Thr
25	Phe	Ala	Arg 107		Asp	Phe	Leu	Glu 108	_	Leu	Glu	Ala	Leu 108!	Lys 5	Asn	Lys
	Arg	Phe 1090		Asp	Ala	Ile	Pro 109		Ala	Met	Asp	Val		Leu	Lys	Tyr
	Ser 110		Arg	Asn	Arg	Val 1110		Asn	Asn	Val	Trp 1115		Thr	Gly	Val	Gly 112
30			Ser	Phe	Ile 1129	Ser		Gly	Thr	Gly	Thr		Tyr	Gly	Ile 113	Asn
	Val	Gly	Tyr	Asp	Arg		Ile	Lys	Gly 1145	Val		Val	Gly	Gly 1150	Tyr	
35	Ala	Tyr	Gly 1155	Tyr		Gly	Phe	His	Ala		Ile	Thr	Gln 1169	Ser		Ser
<i>J J</i>	Ser	Asn 1170	Val		Val	Gly	Val	Tyr		Arg	Ala	Phe 1180	Ile	Lys	Arg	Ser
		Leu		Met	Ser		Asn	-	Thr	Trp		Tyr		Lys	Thr	
40	118		Ser	Tur	Asn	1190		T.e.11	Ser	Tle	1195		Gln	Ser	ጥ ህ ዮ	1200
	110	71011	J C1	- / -	1205		200	Dea	JCI	1210		71011	022		1215	
				1220)				1225	5		_	_	Tyr 1230)	
45	Met	Phe	Lys 1235	_	Lys	Ser	Val	Ile 1240		Lys	Pro	Gln	Val 1245	Gly	Leu	Ser
	Tyr	Tyr 1250	Tyr		Gly	Leu	Ser 1255	Gly		Arg	Gly	Ile 1260	Met	Asp	Asp	Pro
	Ile	Tyr	Asn	Gln	Phe	Arg	Ala	Asn	Ala	Asp	Pro	Asn	Lys	Lys	Ser	Val
50	1265		Tle	Aen	Dhe	1270		Glu	Sar	λνα	1275		Dhe	Asn	Lvs	1280
					1285	;				1290)				1295	i
	Ser	Tyr	Tyr	Phe 1300		Ile	Ala	Asp	Val 1305		Arg	Asp	Leu	Phe 1310		Asn
55	Ser	Met	Gly 1315		Lys	Met	Val	Arg 1320		Ile	Gly	Asn	Asn 1325	Thr	Leu	Ser

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	Tyr	Arg .		Gly	Gly	Arg	Tyr 1335		Thr	Phe	Ala	Ser 1340	Ile	Ile	Thr	GLY
	Glv	Glu	Ile 2	Arg	Leu	Phe			Phe	Tyr	Val	Asn	Ala	Gly	Ile	Gly
	1345	i				1350					1355					1360
5	Ala	Arg	Phe (Tyr	Lys	Asp	Ile	Asn	Ile	Thr	Gly	Asn	Ile
					1365					1370					1375	1
	Gly	Met		191 1380		Pne										
				1300											,	
10	(2)	INFO	RMAT	ION	FOR	SEQ	ID N	0:13	2:							
		(i)	SEQ				TERI 2 am			le						
							o ac		acic							
15							line									
									-							
		(ii)	MOL	ECUL	E TY	PE:	prot	ein								
		(iii)	HAD	ОТИБ	יידר'ם	.T Y	ES									
20		(111)	nir	OINE	11101											
_0		(vi)	ORI													
			(A	OR	GANI	SM:	Heli	.coba	cter	pyl	.ori					
		(\	E1 E1 B	orm to												
25		(1X)	FEA			ŒY:	misc	: fea	ature	.						
23							2									
		(xi)	SEQ	UENC	E DE	ESCRI	PTIC)N: 5	SEQ :	LD NO):132	::				
30	Met.	Lys	Lvs	Ile	Gly	Leu	Ser	Leu	Cys	Leu	Val	Leu	Ser	Leu	Gly	Phe
	1				5					10					15	
	Leu	Lys	Ala		Glu	Val	Ser	Ala		Glu	Ile	Ala	Asp		Phe	Tyr
	•	Leu	7	20	Tue	Glu	Dro	Lve	25 Met	T.VS	Tle	Asn	His	30 Thr	Lvs	Gly
35	Lys	Leu	35	ALA	пуз	GIU	110	40		_			45		- 4	•
55	Phe	Cys		Lys	Gly	Val	Phe	Leu	Pro	Asn	Pro	Gln	Ala	Arg	Glu	Asp
		50					55		•			60				
		Glu	Val	Pro	Leu		Asn	Glu	Lys	GLu	11e 75	Pro	Ala	ser	vai	80
40	65 Tur	Sar	T. (2) 1	Glv	Glv	70 Val	Ala	Met	Asp	Asp	_	Ser	Lys	Val	Arg	Gly
+0	ıyı	261	Deu	Gry	85					90	4		•		95	
	Met	Ala	Leu	Lys	Leu	Glu	Asn	Gln	Asn	Ala	Ser	Trp	Thr		Val	Met
				100					105		5	a 1	G 1	110	ת 1 ת	Gln
A =	Leu	Asn		Glu	Ile	Asn	Phe	A1a 120	Lys	Asn	Pro	GIU	125	Pne	Ala	Gln
45	Dhe	Phe	115 Glu	Met	Δτα	Leu	Pro		Asn	Glv	Lvs	Val		Glu	Ala	Arg
	PHE	130	GIU		****		135	1 -		2		140	-			
	Ile	Lys	Lys	Leu	Tyr	Glu	Glu	Val	Pro	Ser	Tyr	Arg	Asn	Phe	Ala	Ala
=	145					150		_	_	_	155		•	ml	D=0	160
50	Tyr	Met	Lys	Thr		Gly	Ile	Ser	ser	Ser	val	ALA	ASN	Thr	175	Tyr
	TT~	ge ~	Val.	Hie	165 Ala	Phe	Lvs	Phe	Lvs		Lvs	Lvs	Glu	Lys		Leu
	TAT	JET	A CT T	180	44±U		-,-		185		_1_	-1-		190		
	Pro	Ala	Arg		Lys	Phe	Val	Pro	Lys	Glu	Gly	Val		Tyr	Leu	Asn
55			195					200					205			

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pro Gln Glu Leu Lys Gln Lys Asp Ser Asn Tyr Leu Leu Ser Ser Phe 215 Gln Gln His Leu Lys Asn Lys Pro Ile Glu Tyr Gln Met Tyr Leu Val 230 235 Phe Ala Asn Gln Asn Asp Ala Thr Asn Asp Thr Thr Ala Leu Trp Lys 5 245 250 Gly Ser Ile Arg Asn Tyr 260 10 (2) INFORMATION FOR SEQ ID NO:133: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 246 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: protein (iii) HYPOTHETICAL: YES 20 (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori (ix) FEATURE: 25 (A) NAME/KEY: misc_feature (B) LOCATION 1...246 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:133: 30 Met Lys Gln Phe Lys Lys Lys Pro Lys Lys Ile Lys Arg Ser His Gln Asn Gln Lys Thr Ile Leu Lys Arg Pro Leu Trp Leu Met Pro Leu Leu 25 Ile Gly Gly Phe Ala Ser Gly Val Tyr Ala Asp Gly Thr Asp Ile Leu 35 40 Gly Leu Ser Trp Gly Glu Lys Ser Gln Lys Val Cys Val His Arg Pro 55 Trp Tyr Ala Ile Trp Ser Cys Asp Lys Trp Glu Glu Lys Thr Gln Gln 75 40 Phe Thr Gly Asn Gln Leu Ile Thr Lys Thr Trp Ala Gly Gly Asn Ala 90 Ala Asn Tyr Tyr His Ser Gln Asn Asn Gln Asp Ile Thr Ala Asn Leu 105 Lys Asn Asp Asn Gly Thr Tyr Phe Leu Ser Gly Leu Tyr Asn Tyr Thr 45 . 120 Gly Gly Glu Tyr Asn Gly Gly Asn Leu Asp Ile Glu Leu Gly Ser Asn 135 Ala Thr Phe Asn Leu Gly Ala Ser Ser Gly Asn Ser Phe Thr Ser Trp 150 155 50 Tyr Pro Asn Gly His Thr Asp Val Thr Phe Ser Ala Gly Thr Ile Asn 170 Val Asn Asn Ser Val Glu Val Gly Asn Arg Val Gly Ser Gly Ala Gly 185 Thr His Thr Gly Thr Ala Thr Leu Asn Leu Asn Ala Asn Lys Val Thr 55 200

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Ile Asn Ser Asn Ile Ser Ala Tyr Lys Thr Ser Gln Val Asn Val Gly 215 Asn Ala Asn Ser Val Ile Thr Ile Asn Ser Val Ser Leu Asn Gly Glu 230 Tyr Leu Gln Phe Phe Ser (2) INFORMATION FOR SEQ ID NO:134: (i) SEQUENCE CHARACTERISTICS: 10 (A) LENGTH: 245 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (ii) MOLECULE TYPE: protein 15 (iii) HYPOTHETICAL: YES (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori 20 (ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1...245 25 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:134: Met Ile Lys Lys Thr Leu Ala Ser Val Leu Leu Gly Leu Ser Leu Met 10 Ser Val Leu Asn Ala Lys Glu Cys Val Ser Pro Ile Thr Arg Ser Val 30 25 Lys Tyr His Gln Gln Ser Ala Glu Ile Arg Ala Leu Gln Leu Gln Ser 40 Tyr Lys Met Ala Lys Met Ala Leu Asp Asn Asn Leu Lys Leu Val Lys 35 Asp Lys Lys Pro Ala Val Ile Leu Asp Leu Asp Glu Thr Val Leu Asn 70 Thr Phe Asp Tyr Ala Gly Tyr Leu Val Lys Asn Cys Ile Lys Tyr Thr Pro Glu Thr Trp Asp Lys Phe Glu Lys Glu Gly Ser Leu Thr Leu Ile 40 105 Pro Gly Ala Leu Asp Phe Leu Glu Tyr Ala Asn Ser Lys Gly Val Lys 125 120 Ile Phe Tyr Ile Ser Asn Arg Thr Gln Lys Asn Lys Ala Phe Thr Leu 140 45 135 Lys Thr Leu Lys Ser Phe Lys Leu Pro Gln Val Ser Glu Glu Ser Val 155 150 Leu Leu Lys Glu Lys Gly Lys Pro Lys Ala Val Arg Arg Glu Leu Val 170 Ala Lys Asp Tyr Ala Ile Val Leu Gln Val Gly Asp Thr Leu His Asp 50 185 180 Phe Asp Ala Ile Phe Ala Lys Asp Ala Lys Asn Ser Gln Glu Gln Gln 200

Ala Lys Val Leu Gln Asn Ala Gln Lys Phe Gly Thr Glu Trp Ile Ile

215

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Leu Pro Asn Ser Leu Tyr Gly Thr Trp Glu Asp Gly Pro Ile Lys Ala 230 235 Trp Gln Asn Lys Lys 5 (2) INFORMATION FOR SEQ ID NO:135: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 288 amino acids 10 (B) TYPE: amino acid (D) TOPOLOGY: linear (ii) MOLECULE TYPE: protein 15 (iii) HYPOTHETICAL: YES (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori 20 (ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1...288 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:135: 25 Leu Trp Cys Leu Lys Thr Pro Ile Ile Gly His Gly Met Lys Lys Ala Lys Val Phe Trp Cys Cys Phe Lys Met Ile Arg Trp Leu Tyr Leu Ala Val Phe Phe Leu Leu Ser Val Ser Asp Ala Lys Glu Ile Ala Met 30 40 Gln Arg Phe Asp Lys Gln Asn His Lys Ile Phe Glu Ile Leu Ala Asp Lys Val Ser Ala Lys Asp Asn Val Ile Thr Ala Ser Gly Asn Ala Ile 35 70 75 Leu Leu Asn Tyr Asp Val Tyr Ile Leu Ala Asp Lys Val Arg Tyr Asp 90 Thr Lys Thr Lys Glu Ala Leu Leu Glu Gly Asn Ile Lys Val Tyr Arg 105 110 40 Gly Glu Gly Leu Leu Val Lys Thr Asp Tyr Val Lys Leu Ser Leu Asn 120 Glu Lys Tyr Glu Ile Ile Phe Pro Phe Tyr Val Gln Asp Ser Val Ser 135 140 Gly Ile Trp Val Ser Ala Asp Ile Ala Ser Gly Lys Asp Gln Lys Tyr 45 150 . 155 Lys Ile Lys Asn Met Ser Ala Ser Gly Cys Ser Ile Asp Asn Pro Ile 165 170 Trp His Val Asn Ala Thr Ser Gly Ser Phe Asn Met Gln Lys Ser His 185 50 Leu Ser Met Trp Asn Pro Lys Ile Tyr Val Gly Asp Ile Pro Val Leu 200 Tyr Leu Pro Tyr Ile Phe Met Ser Thr Ser Asn Lys Arg Thr Thr Gly 215 Phe Leu Tyr Pro Glu Phe Gly Thr Ser Asn Leu Asp Gly Phe Ile Tyr 55 230 235

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Leu Gln Pro Phe Tyr Leu Ala Pro Lys Asn Ser Trp Asp Met Thr Phe 250 Thr Pro Gln Ile Arg Tyr Lys Arg Gly Phe Gly Leu Asn Phe Glu Ala 265 270 5 Arg Tyr Ile Asn Ser Lys Thr Gln Val Phe Ile Gln Cys Ala Leu Phe 280 (2) INFORMATION FOR SEQ ID NO:136: 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 128 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: protein (iii) HYPOTHETICAL: YES (vi) ORIGINAL SOURCE: 20 (A) ORGANISM: Helicobacter pylori (ix) FEATURE: (A) NAME/KEY: misc feature (B) LOCATION 1...128 25 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:136: Leu Met Phe Lys Lys Met Cys Leu Ser Leu Leu Met Ile Ser Gly Val 10 30 Cys Val Gly Ala Lys Asp Leu Asp Phe Lys Leu Asp Tyr Arg Ala Thr 25 Gly Gly Lys Phe Met Gly Lys Met Thr Asp Ser Ser Leu Leu Ser Ile 40 Thr Ser Met Asn Asp Glu Pro Val Val Ile Lys Asn Leu Ile Val Asn 35 Arg Gly Asn Ser Cys Glu Ala Thr Lys Lys Val Glu Pro Lys Phe Gly 70 75 Asp Lys Phe Lys Lys Glu Lys Leu Phe Asp His Glu Leu Lys Tyr Ser 85 90 40 Gln Gln Ile Phe Tyr Arg Leu Asp Cys Lys Pro Asn Gln Leu Leu Glu 105 Val Lys Ile Ile Thr Asp Lys Gly Glu Tyr Tyr His Lys Phe Ser Lys 120 45 (2) INFORMATION FOR SEQ ID NO:137: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 169 amino acids (B) TYPE: amino acid 50 (D) TOPOLOGY: linear

(iii) HYPOTHETICAL: YES 55

(ii) MOLECULE TYPE: protein

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(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori (ix) FEATURE: 5 (A) NAME/KEY: misc_feature (B) LOCATION 1...169 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:137: Met Gln Ala Leu Lys Ser Leu Leu Glu Val Ile Thr Lys Leu Gln Asn 10 Leu Gly Gly Tyr Leu Met His Ile Ala Ile Phe Ile Ile Phe Ile Trp Ile Gly Gly Leu Lys Phe Val Pro Tyr Glu Ala Glu Gly Ile Ala Pro 15 40 Phe Val Ala Asn Ser Pro Phe Phe Ser Phe Met Tyr Lys Phe Glu Lys 55 Pro Ala Tyr Lys Gln His Lys Met Ser Glu Ser Gln Ser Met Gln Glu 70 75 20 Glu Met Gln Asp Asn Pro Lys Ile Val Glu Asn Lys Glu Trp His Lys 85 90 Glu Asn Arg Thr Tyr Leu Val Ala Glu Gly Leu Gly Ile Thr Ile Met 105 Ile Leu Gly Ile Leu Val Leu Leu Gly Leu Trp Met Pro Leu Met Gly 25 120 Val Val Gly Gly Leu Leu Val Ala Gly Met Thr Ile Thr Thr Leu Phe 135 140 Phe Phe Ile His Asn Ala Arg Ser Val Cys Gln Ser Ala Phe Pro Met 150 30 Ala Phe Trp Gly Trp Lys Ala Ser Gly 165 (2) INFORMATION FOR SEQ ID NO:138: 35 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 487 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear 40 (ii) MOLECULE TYPE: protein (iii) HYPOTHETICAL: YES (vi) ORIGINAL SOURCE: 45 (A) ORGANISM: Helicobacter pylori (ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1...487 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:138: Met Ile Glu Trp Met Gln Asn His Arg Lys Tyr Leu Val Val Thr Ile

Trp Ile Ser Thr Ile Ala Phe Ile Ala Ala Gly Met Ile Gly Trp Gly

			•	20					25					30		
			35	Phe				40					45	Val		
5		50					55					60		Arg		
	65					70					75			Thr		80
					85					90				Ser	95	
10	Asn			100					105					110		
			115					120					125	Val		
15	_	130					135					140		Leu		
	145					150					155			Arg		160
• •					165					170				Thr	175	
20				180					185					Lys 190		
			195					200					205	Asn Lys		
25		210			•		215					220		Leu		
	225					230					235			Lys		240
30					245					250					255	Glu
30				260					265					270		Leu
			275					280					285			Thr
35		290					295					300				Gln
	305					310					315					320 Phe
40					325					330					335	Glu
				340					345					350		Thr
			355					360					365			Leu
45		370					375					380				Gly
	385					390					395				Ile	Asn
50					405					410				Ile	415 Gly	Asn
				420					425				Asn	430 His		Phe
			435					440				Val	Asn Asn			Lys
55		450					455					460				

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Thr Asp Phe Phe Asp Lys Ala Leu Ile Glu Glu Leu Lys Lys Arg Tyr 470 475 Lys Ile Val Lys Tyr Ile Gln 485 5 (2) INFORMATION FOR SEQ ID NO:139: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 142 amino acids 10 (B) TYPE: amino acid (D) TOPOLOGY: linear (ii) MOLECULE TYPE: protein 15 (iii) HYPOTHETICAL: YES (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori 20 (ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1...142 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:139: 25 Met Lys Thr Asn Phe Tyr Lys Ile Lys Leu Leu Phe Ala Trp Cys Leu Ile Ile Gly Met Phe Asn Ala Pro Leu Asn Ala Asp Gln Asn Thr Asp 25 30 Ile Lys Asp Ile Ser Pro Glu Asp Met Ala Leu Asn Ser Val Gly Leu 40 Val Ser Arg Asp Gln Leu Lys Ile Glu Ile Pro Lys Glu Thr Leu Glu 55 Gln Lys Val Ala Ile Leu Asn Asp Tyr Asn Asp Lys Asn Val Asn Ile 35 70 75 Lys Phe Asp Asp Ile Ser Leu Gly Ser Phe Gln Pro Asn Asp Asn Leu 90 85 Gly Ile Asn Ala Met Trp Gly Ile Gln Asn Leu Leu Met Ser Gln Met 105 40 Met Ser Asn Tyr Gly Pro Asn Asn Ser Phe Met Tyr Gly Tyr Ala Pro 120 Thr Tyr Ser Asp Ser Ser Phe Leu Pro Pro Ile Leu Gly Tyr 135 45 (2) INFORMATION FOR SEQ ID NO:140: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 208 amino acids (B) TYPE: amino acid 50 (D) TOPOLOGY: linear (ii) MOLECULE TYPE: protein

(iii) HYPOTHETICAL: YES

INCOCCIO: JMO GRIPTIZANI I

(vi) ORIGINAL SOURCE:

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(A) ORGANISM: Helicobacter pylori
         (ix) FEATURE:
 5
               (A) NAME/KEY: misc_feature
               (B) LOCATION 1...208
         (xi) SEQUENCE DESCRIPTION: SEQ ID NO:140:
     Leu Ile Asn Asn Asn Asn Asn Lys Lys Leu Arg Gly Phe Phe Leu
10
     Lys Val Leu Leu Ser Leu Val Val Phe Ser Ser Tyr Gly Ser Ala Asn
                                     25
     Asp Asp Lys Glu Ala Lys Lys Glu Ala Leu Glu Lys Glu Lys Asn Thr
15
                                 40
     Pro Asn Gly Leu Val Tyr Thr Asn Leu Asp Phe Asp Ser Phe Lys Ala
                                                 60
     Thr Ile Lys Asn Leu Lys Asp Lys Lys Val Thr Phe Lys Glu Val Asn
                                             75
     Pro Asp Ile Ile Lys Asp Glu Val Phe Asp Phe Val Ile Val Asn Arg
20
                                         90
     Val Leu Lys Lys Ile Lys Asp Leu Lys His Tyr Asp Pro Val Ile Glu
                                     105
     Lys Ile Phe Asp Glu Lys Gly Lys Glu Met Gly Leu Asn Val Glu Leu
25
                                 120
     Gln Ile Asn Pro Glu Val Lys Asp Phe Phe Thr Phe Lys Ser Ile Ser
                             135
     Thr Thr Asn Lys Gln Arg Cys Phe Leu Ser Leu His Gly Glu Thr Arg
                                             155
                         150
     Glu Ile Leu Cys Asp Asp Lys Leu Tyr Asn Val Leu Leu Ala Val Phe
30
                                         170
     Asn Ser Tyr Asp Pro Asn Asp Leu Leu Lys His Ile Ser Thr Ile Glu
                                    185
     Ser Leu Lys Lys Ile Phe Tyr Thr Ile Thr Cys Glu Ala Val Tyr Leu
35
                                200
     (2) INFORMATION FOR SEQ ID NO:141:
          (i) SEQUENCE CHARACTERISTICS:
               (A) LENGTH: 245 amino acids
40
               (B) TYPE: amino acid
               (D) TOPOLOGY: linear
         (ii) MOLECULE TYPE: protein
45
        (iii) HYPOTHETICAL: YES
         (vi) ORIGINAL SOURCE:
               (A) ORGANISM: Helicobacter pylori
50
         (ix) FEATURE:
               (A) NAME/KEY: misc_feature
               (B) LOCATION 1...245
         (xi) SEQUENCE DESCRIPTION: SEQ ID NO:141:
55
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Met Ala Gly Thr Gln Ala Ile Tyr Glu Ser Ser Ala Gly Phe Leu Ser Gln Val Ser Ser Ile Ile Ser Ser Thr Ser Gly Val Ala Gly Pro 5 25 Phe Ala Gly Ile Val Ala Gly Ala Met Thr Ala Ala Ile Ile Pro Ile 40 Val Val Gly Phe Thr Asn Pro Gln Met Thr Ala Ile Met Thr Gln Tyr 55 10 Asn Gln Ser Ile Ala Glu Ala Val Ser Val Pro Met Lys Ala Ala Asn 70 Gln Gln Tyr Asn Gln Leu Tyr Gln Gly Phe Asn Asp Gln Ser Met Ala Val Gly Asn Asn Ile Leu Asn Ile Ser Lys Leu Thr Gly Glu Phe Asn 15 105 Ala Glm Gly Asn Thr Glm Ser Ala Glm Ile Ser Ala Val Asn Ser Glm 120 Ile Ala Ser Ile Leu Ala Ser Asn Thr Thr Pro Lys Asn Pro Ser Ala 135 140 20 Ile Glu Ala Tyr Ala Thr Asn Gln Ile Ala Val Pro Ser Val Pro Thr 150 155 Thr Val Glu Met Met Ser Gly Ile Leu Gly Asn Ile Thr Ser Ala Ala 165 170 Pro Lys Tyr Ala Leu Ala Leu Gln Glu Gln Leu Arg Ser Gln Ala Ser 25 185 Asn Ser Ser Met Asn Asp Thr Ala Asp Ser Leu Asp Ser Cys Thr Ala 200 Leu Gly Ala Leu Val Gly Ser Ser Lys Val Phe Phe Ser Cys Met Gln Ile Ser Met Thr Pro Met Ser Val Ser Met Pro Thr Val Met Pro Asn 30 230 235 Thr Ser Gly Cys His 35 (2) INFORMATION FOR SEQ ID NO:142: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 367 amino acids (B) TYPE: amino acid 40 (D) TOPOLOGY: linear (ii) MOLECULE TYPE: protein (iii) HYPOTHETICAL: YES 45 (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori (ix) FEATURE: 50 (A) NAME/KEY: misc_feature (B) LOCATION 1...367

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:142:

Met Ile Lys Ser Val Glu Ile Glu Asn Tyr Lys Asn Phe Glu His Leu

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	1				5					10					15	
		Met	Glu	Asn 20	Phe	Lys	Leu	Ile	Asn 25	Phe	Phe	Thr	Gly	Gln 30	Asn	Asp
5	Ala	Gly	Lys 35	Thr	Asn	Leu	Leu	Glu 40	Ala	Leu	Tyr	Thr	Asn 45	Thr	Gly	Leu
	-	50					55					60				
	65					Arg 70					75					80
10				_	85	Gly				90					95	
				100		Thr			105					110		
15			115			Asp		120					125			
		130				Thr	135					140				
20	145					Ser 150					155					160
20					165	Ile				170					175	
	_	_		180		Met			185					190		
25			195			Glu		200					205			
		210				Gln	215					220				
20	225					Val 230 Lys					235					240
30					245	Met				250					255	
		_		260		Glu			265					270		
35	-		275			Ala		280					285			
		290			_	Thr	295					300				
40	305					310					315					320
40					325	Asn				330					335	
				340		Glu			345					350		- y -
45	Ser	Met	Leu 355	GLU	гуѕ	Ala	Leu	1yr 360	Arg	GTÅ	met	GIU	365	Arg	GTÅ	

(2) INFORMATION FOR SEQ ID NO:143:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 409 amino acids

(B) TYPE: amino acid

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

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(iii) HYPOTHETICAL: YES
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(vi) ORIGINAL SOURCE:

(A) ORGANISM: Helicobacter pylori

5

(ix) FEATURE:

(A) NAME/KEY: misc feature

(B) LOCATION 1...409

10 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:143:

Met Ser Leu Ile Arg Val Asn Gly Glu Ala Phe Lys Leu Ser Leu Glu Ser Leu Glu Glu Asp Pro Phe Glu Thr Lys Glu Thr Leu Glu Thr Leu 15 30 25 Glu Thr Leu Ile Lys Gln Thr Ser Val Val Leu Leu Ala Ala Gly Glu 40 Ser Lys Arg Phe Ser Arg Ala Ile Lys Lys Gln Trp Leu Arg Ser His 20 His Thr Pro Leu Trp Leu Ser Val Tyr Glu Ser Phe Lys Glu Ala Leu Asp Phe Lys Glu Val Ile Leu Val Val Ser Glu Leu Asp Tyr Val Tyr Ile Gln Arg His Tyr Pro Lys Ile Lys Leu Val Lys Gly Gly Ala Ser 25 105 Arg Gln Glu Ser Val Arg Asn Ala Leu Lys Val Ile Asp Ser Thr Tyr 120 Thr Ile Thr Ser Asp Val Ala Arg Gly Leu Ala Asn Met Glu Ala Leu 135 140 Lys Ser Leu Phe Leu Thr Leu Gln Gln Thr Ser His Tyr Cys Ile Ala 30 150 Pro Tyr Leu Pro Cys Tyr Asp Thr Ala Ile Tyr Tyr Asn Glu Ala Leu 165 170 Asp Arg Glu Ala Ile Lys Leu Ile Gln Thr Pro Gln Leu Ser His Thr 35 185 Lys Thr Leu Gln Ser Ala Leu Asn Gln Gly Gly Phe Lys Asp Glu Ser 200 Ser Ala Ile Leu Gln Ala Phe Pro Asn Ser Val Ser Tyr Ile Glu Gly 215 220 40 Ser Lys Asp Leu His Lys Leu Thr Thr Ser Gly Asp Leu Lys Phe Phe 230 235 Thr Pro Phe Phe Asn Pro Ala Lys Asp Thr Phe Ile Gly Met Gly Phe 250 Asp Thr His Ala Phe Ile Lys Asp Lys Pro Met Val Leu Gly Gly Val 45 265 Val Leu Asp Cys Glu Phe Gly Leu Lys Ala His Ser Asp Gly Asp Ala 280 Leu Leu His Ala Val Ile Asp Ala Ile Leu Gly Ala Ile Lys Gly Gly 295 50 Asp Ile Gly Glu Trp Phe Pro Asp Asn Asp Pro Lys Tyr Lys Asn Ala 310 315 Ser Ser Lys Glu Leu Leu Lys Ile Val Leu Asp Phe Ser Gln Ser Ile 325 330 Gly Phe Glu Leu Glu Met Gly Ala Thr Ile Phe Ser Glu Ile Pro 55 345

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Lys Ile Thr Pro Tyr Lys Pro Ala Ile Leu Glu Asn Leu Ser Gln Leu 360 Leu Gly Leu Glu Lys Ser Gln Ile Ser Leu Lys Ala Thr Thr Met Glu 375 Lys Met Gly Phe Ile Gly Lys Gln Glu Gly Leu Leu Val Gln Ala His 390 395 Val Ser Met Arg Tyr Lys Gln Lys Leu 405 (2) INFORMATION FOR SEQ ID NO:144: 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 270 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: protein (iii) HYPOTHETICAL: YES 20 (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori (ix) FEATURE: (A) NAME/KEY: misc_feature 25 (B) LOCATION 1...270 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:144: Met Lys Lys Phe Val Ala Leu Gly Leu Leu Ser Ala Val Leu Ser Ser 30 Ser Leu Leu Ala Glu Gly Asp Gly Val Tyr Ile Gly Thr Asn Tyr Gln 25 Leu Gly Gln Ala Arg Leu Asn Ser Asn Ile Tyr Asn Thr Gly Asp Cys 35 Thr Gly Ser Val Val Gly Cys Pro Pro Gly Leu Thr Ala Asn Lys His 55 Asn Pro Gly Gly Thr Asn Ile Asn Trp His Ser Lys Tyr Ala Asn Gly Ala Leu Asn Gly Phe Gly Leu Asn Val Gly Tyr Lys Lys Phe Phe Gln 40 90 85 Phe Lys Ser Leu Asp Met Thr Ser Lys Trp Phe Gly Phe Arg Val Tyr 105 Gly Leu Phe Asp Tyr Gly His Ala Asp Leu Gly Lys Gln Val Tyr Ala 45 120 Pro Asn Lys Ile Gln Leu Asp Met Val Ser Trp Gly Val Gly Ser Asp 140 135 Leu Leu Ala Asp Ile Ile Asp Lys Asp Asn Ala Ser Phe Gly Ile Phe 155 150 Gly Gly Val Ala Ile Gly Gly Asn Thr Trp Lys Ser Ser Ala Ala Asn 50 170 Tyr Trp Lys Glu Gln Ile Ile Glu Ala Lys Gly Pro Asp Val Cys Thr 185 Pro Thr Tyr Cys Asn Pro Asn Ala Pro Tyr Ser Thr Asn Thr Ser Thr 200 55

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Val Ala Phe Gln Val Trp Leu Asn Phe Gly Val Arg Ala Asn Ile Tyr 215 Lys His Asn Gly Val Glu Phe Gly Val Arg Val Pro Leu Leu Ile Asn 230 235 Lys Phe Leu Ser Ala Gly Pro Asn Ala Thr Asn Leu Tyr Tyr His Leu 5 250 245 Lys Arg Asp Tyr Ser Leu Tyr Leu Gly Tyr Asn Tyr Thr Phe 260 . 265 (2) INFORMATION FOR SEQ ID NO:145: 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 438 amino acids (B) TYPE: amino acid 15 (D) TOPOLOGY: linear (ii) MOLECULE TYPE: protein (iii) HYPOTHETICAL: YES 20 (vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori (ix) FEATURE: 25 (A) NAME/KEY: misc feature (B) LOCATION 1...438 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:145: 30 Met Ala Tyr Lys Pro Asn Lys Lys Leu Lys Glu Leu Arg Glu Gln Pro Asn Leu Phe Ser Ile Leu Asp Lys Gly Asp Val Ala Thr Asn Asn Pro Val Glu Glu Ser Asp Lys Ala Asn Lys Ile Gln Glu Pro Leu Pro 35 40 Tyr Val Val Lys Thr Gln Ile Asn Lys Ala Ser Met Ile Ser Arg Asp 55 Pro Ile Glu Trp Ala Lys Tyr Leu Ser Phe Glu Lys Arg Val Tyr Lys 75 40 Asp Asn Ser Lys Glu Asp Val Asn Phe Phe Ala Asn Gly Glu Ile Lys Glu Ser Ser Arg Val Tyr Glu Ala Asn Lys Glu Gly Phe Glu Arg Arg 105 Ile Thr Lys Arg Tyr Asp Leu Ile Asp Arg Asn Ile Asp Arg Asn Arg 45 120 Glu Phe Phe Ile Lys Glu Ile Glu Ile Leu Thr His Thr Asn Ser Leu 135 Lys Glu Leu Lys Glu Gln Gly Leu Glu Ile Gln Leu Thr His His Asn 150 155 50 Glu Thr His Lys Lys Ala Leu Glu Asn Gly Asn Glu Ile Val Lys Glu 170 Tyr Asp His Leu Lys Asp Ile Tyr Gln Glu Val Glu Arg Thr Lys Asp 185 Gly Gly Leu Val Arg Glu Ile Ile Pro Ser Ile Ser Ser Ala Glu Tyr 55 200

		210					215					220	Asn			
	225					230					235		Lys			240
5	Glu	Leu	Ala	Lys	Glu 245	Val	His	Ile	Leu	Ile 250	Leu	Glu	Gln	Gln	Leu 255	Leu
	Ser	Ala	Thr	Asn 260		Tyr	Ser	Trp	Ile 265	Asp	Lys	Asp	Asp	Asn 270	Ala	Asn
10	Phe	Ala	Trp 275	Lys	Met	His	Àrg	Leu 280		Asn	Glu	Asn	Lys 285	Leu	Lys	Glu
10	Asn	His 290	Leu	Ser	Ala	Asn	Asn 295		Asn	Lys	Ile	Lys 300	Gln	Phe	Phe	Phe
	Asn	Asn	Gly	Ser	Ile	Leu	Gly	Trp	Thr	Lys	Glu	Glu	Gln	Ser	Ala	Ile
	305					310					315					320
15					325					330			Leu -		335	
				340					345				Tyr	350		
20	Tyr	Val	Asn 355	Gly	Asp	Gly	Asn	Lys 360	Arg	Glu	Ile	Lys	Pro 365	Pne	гÀг	GIU
20	Ile	Leu 370	Arg	Asp	Thr	Asn	Asn 375		Glu	Lys	Ala	Tyr 380	Lys	Glu	Arg	Tyr
	Asp	Lys	Leu	Val	Ser	Leu		Ala	Ala	Ile	Ile	Gln	Ala	Lys	Glu	Gly
	385					390					395					400
25					405					410			Asn		415	
	Asn	Thr	Ile	Glu 420		Asn	Thr	Ser	Asn 425		Ile	Ile	Gln	430	Asn	Asp
30	Asn	Ile	Ile 435		Gln	Ile										
	(2)	INF	ORMA	TION	FOR	SEQ	ID :	NO:1	46:							
35		(i	(A) L B) T	ENGT YPE :	HARA H: 2 ami OGY:	15 a no a	mino cid		ds						
40		(ii) MO	LECU	LE T	YPE:	pro	tein								
40		(iii) HY	РОТН	ETIC	AL:	YES									
45		(vi) OR (IGIN A) O	AL S RGAN	OURC	E: Hel	icob	acte	r py	lori					
73		(ix	(AME/	KEY:			atur	e				٠		
50		(xi) SE	QUEN	CE I	ESCR	IPTI	ON:	SEQ	ID N	10:14	6:				
	Met 1	Gln	Ala	Lev	Lys 5	Ser	Leu	Leu	Glu	Val	. Ile	Thr	Lys	Leu	Gln 15	Ası
55	Leu	Gly	Gly	7 Tyr 20	Lev	Met	His	Ile	Ala 25	ı Ile	Phe	: Ile	lle	Phe 30	Ile	Trj

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	Ile	Gly	Gly 35	Leu	Lys	Phe	Val	Pro 40	Tyr	Glu	Ala	Glu	Gly 45	Ile	Ala	Pro
	Phe	Val 50	Ala	Asn	Ser	Pro	Phe 55	Phe	Ser	Phe	Met	Tyr 60	Lys	Phe	Glu	Lys
5	Pro 65	Ala	Tyr	Lys	Gln	His 70	Lys	Met	Ser	Glu	Ser 75	Gln	Ser	Met	Gln	Glu 80
	Glu	Met	Gln	Asp	Asn 85	Pro	Lys	Ile	Val	Glu 90	Asn	Lys	Glu	Trp	His 95	Lys
10	Glu	Asn	Arg	Thr 100	Tyr	Leu	Val	Ala	Glu 105	Gly	Leu	Gly	Ile	Thr 110	Ile	Met
	Ile	Leu	Gly 115	Ile	Leu	Val	Leu	Leu 120	Gly	Leu	Trp	Met	Pro 125	Leu	Met	Gly
	Val	Val 130	Gly	Gly	Leu	Leu	Val 135	Ala	Gly	Met	Thr	Ile 140	Thr	Thr	Leu	Ser
15	Phe 145	Leu	Phe	Thr	Thr	Pro	Glu	Val	Phe	Val	Asn 155		His	Phe	Pro	
						Arg							Ala	Leu	Phe 175	160 Ala
20	Gly	Gly	Leu	Phe 180	Val	Ala	Gly	Phe	Asp 185	Ala	Lys	Arg	Tyr	Leu 190	Glu	Gly
	Lys	Gly	Phe 195	Cys	Leu	Met	Asp	Arg 200	Ser	Ser	Val	Gly	Ile 205	Lys	Thr	Lys
25	Cys	Ser 210	Ser	Gly	Cys	Cys	Ser 215									
25	(2)	INFO	ORMAT	rion	FOR	SEQ	ID 1	10:14	17:							
30		(i)	(<i>I</i>	A) LE B) TY C) ST	ENGTH (PE: TRANI	IARAC I: 20 nucl EDNE	bas eic SS:	se pa ació doub	irs l ole							
35	÷ ,					PE:		(ger	omic	:)				•		
	·		ANT				.0									
40		(vi)				URCE		.coba	cter	pyl	.ori					
45		(ix)) NA	ME/K	EY:		_	ture	:						
		(xi)	SEÇ	UENC	E DE	SCRI	PTIC	N: S	EQ I	D NO	:147	' :				
50	TATA	CCAT	GG I	'GGGC	GCTA	A										
	(2)	INFO	RMAT	NOI	FOR	SEQ	ID N	10:14	8:							
		(i)				ARAC										
55						nucl		-								

55

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		(C) STRANDEDNESS: double (D) TOPOLOGY: circular	•
_	(ii)	MOLECULE TYPE: DNA (genomic)	
5	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
10	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
15		FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 123	
		SEQUENCE DESCRIPTION: SEQ ID NO:148:	22
20	ATGAATTCG	A GTAAGGATTT TTG	23
	(2) INFOR	MATION FOR SEQ ID NO:149:	-
25	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 22 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
30		MOLECULE TYPE: DNA (genomic)	
		HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
35	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
40	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 122	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:149:	
45	TTAACCATO	GG TGAAAAGCGA TA	22
45	(2) INFO	RMATION FOR SEQ ID NO:150:	
50	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
55	(ii)	MOLECULE TYPE: DNA (genomic)	

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	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
5	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
10	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 123	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:150:	
15	TAGAATTC	GC ATAACGATCA ATC	23
	-('2') INFO	RMATION FOR SEQ ID NO:151:	
20	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 22 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	·
25	(ii)	MOLECULE TYPE: DNA (genomic)	
25	(iii)	HYPOTHETICAL: NO	,
	(iv)	ANTI-SENSE: NO	
30	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
35	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 122	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:151:	
40	ATATCCAT	GG TGAGTTTGAT GA	22
70	(2) INFO	RMATION FOR SEQ ID NO:152:	
45	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 25 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
50	(ii)	MOLECULE TYPE: DNA (genomic)	
50	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
55	(3ri)	OPIGINAL SOURCE.	

		(A) ORGANISM: Helicobacter pylori	
5	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 125	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:152:	
	ATGAATTCA	A TTTTTTATTT TGCCA	25
10	(2) INFOR	MATION FOR SEQ ID NO:153:	
15	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 21 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
20	(ii)	MOLECULE TYPE: DNA (genomic)	
20	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
25	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
30	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 121	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:153:	
25	AATTCCAT	GG TGGGGGCTAT G	21
35	(2) INFO	RMATION FOR SEQ ID NO:154:	
40	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
4.5	(ii)	MOLECULE TYPE: DNA (genomic)	
45	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
50	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
55	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 123	

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	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:154:	
5	ATGAATTCTC GATAGCCAAA ATC	23
3	(2) INFORMATION FOR SEQ ID NO:155:	
10	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 25 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
15	(ii) MOLECULE TYPE: DNA (genomic)	
13	(iii-)- HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
20	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
25	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 125</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:155:	
30	AATTCCATGG TGCATAACTT CCATT	25
	(2) INFORMATION FOR SEQ ID NO:156:	•
35	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 25 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
40	(ii) MOLECULE TYPE: DNA (genomic)	
40	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
45	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
50	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 125</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:156:	
55	AAGAATTCTC TAGCATCCAA ATGGA	25

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	(2) INFORMATION FOR SEQ ID NO:157:	
5	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 24 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
10	(ii) MOLECULE TYPE: DNA (genomic)	
10	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
15	(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
20	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 124</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:157:	
25	ATTTCCATGG TCATGTCTCA TATT	24
25	(2) INFORMATION FOR SEQ ID NO:158:	
30	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
35	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
40	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
45	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 123</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:158:	
50	ATGAATTCCA TCTTTTATTC CAC	23
<i>5</i> 0	(2) INFORMATION FOR SEQ ID NO:159:	
55	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 27 base pairs(B) TYPE: nucleic acid	

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	<pre>(C) STRANDEDNESS: double (D) TOPOLOGY: circular</pre>	
5	(ii) MOLECULE TYPE: DNA (genomic)	
J	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
10	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
15	(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 127	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:159:	
20	AACCATGGTG ATTTTAAGCA TTGAAAG	2'
20	(2) INFORMATION FOR SEQ ID NO:160:	
25	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 28 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: circular	
30	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
35	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
40	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 128</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:160:	
45	AAGAATTCCA CTCAAAATTT TTTAACAG	28
	(2) INFORMATION FOR SEQ ID NO:161:	
50	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 25 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
55	(ii) MOLECULE TYPE: DNA (genomic)	

	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
5	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
10	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 125	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:161:	
15	GATCATCC	AT ATGTTATCTT CTAAT	25
15	(2) INFO	RMATION FOR SEQ ID NO:162:	
20	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
25	(ii)	MOLECULE TYPE: DNA (genomic)	
23	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
30	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
35	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 123	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:162:	
40	TGAATTCA	AC CATTTTAACC CTG	23
40	(2) INFO	RMATION FOR SEQ ID NO:163:	
45	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 27 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
50	(ii)	MOLECULE TYPE: DNA (genomic)	
30	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
55	(vi)	ORIGINAL SOURCE:	

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	(A) ORGANISM: Helicobacter pylori	
5	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 127</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:163:	
••	TATACCATGG TGAAATTTTT TCTTTTA	27
10	(2) INFORMATION FOR SEQ ID NO:164:	
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 25 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	w -
20	(ii) MOLECULE TYPE: DNA (genomic)	٤,
20	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
25	(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
30	(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 125	-
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:164:	
35	AGAATTCAAT TGCGTCTTGT AAAAG	25
33	(2) INFORMATION FOR SEQ ID NO:165:	
40	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 24 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
45	(ii) MOLECULE TYPE: DNA (genomic)	
73	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
50	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
5	(ix) FEATURE: (A) NAME/KEY: misc_feature	

	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:165:	
	TATACCATGG TGATGGACAA ACTC	24
5	(2) INFORMATION FOR SEQ ID NO:166:	
10	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
15	(ii) MOLECULE TYPE: DNA (genomic)	
13	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
20	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
25	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 123</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:166:	
30	ATGAATTCCC ACTTGGGGCG ATA	23
	(2) INFORMATION FOR SEQ ID NO:167:	
35	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 25 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
40	(ii) MOLECULE TYPE: DNA (genomic)	
40	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
45	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
50	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 125</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:167:	
55	TTATGGATCC AAACCAATTA AAACT	25

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	(2) INFORMATION FOR SEQ ID NO:168:	
5	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
10	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
15	(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
	(ix) FEATURE:	
20	(A) NAME/KEY: misc_feature (B) LOCATION 123	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:168:	·
25	TATCTCGAGT TATAGAGAAG GGC	23
23	(2) INFORMATION FOR SEQ ID NO:169:	
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 22 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: circular	
35	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
40	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
45	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 122</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:169:	
- 0	TTAACCATGG TGAAAAGCGA TA	22
50	(2) INFORMATION FOR SEQ ID NO:170:	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 24 base pairs	
55	(B) TYPE: nucleic acid	

		(C) STRANDEDNESS: double (D) TOPOLOGY: circular	
	(ii)	MOLECULE TYPE: DNA (genomic)	
5	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
10	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
15		FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 124 SEQUENCE DESCRIPTION: SEQ ID NO:170:	
			24
20		GC CTCTAAAACT TTAG	
25		SEQUENCE CHARACTERISTICS: (A) LENGTH: 22 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
30		MOLECULE TYPE: DNA (genomic)	
		HYPOTHETICAL: NO	
		ANTI-SENSE: NO	
35	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	٠
40	(ix)	<pre>FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 122</pre>	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:171:	
45	TTAACCAT	GG TGAAAAGCGA TA	22
73	(2) INFO	RMATION FOR SEQ ID NO:172:	
50	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
55	(ii)	MOLECULE TYPE: DNA (genomic)	

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	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
5	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
10	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 123	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:172:	
15	TAGAATTC	GC ATAACGATCA ATC	23
	(2) INFO	RMATION FOR SEQ ID NO:173:	
20	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 22 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
25	(ii)	MOLECULE TYPE: DNA (genomic)	
23	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
30	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
35	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 122	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:173:	
40	ATATCCATO	GG TGAGTTTGAT GA	22
	(2) INFO	RMATION FOR SEQ ID NO:174:	
45	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 25 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
50	(ii)	MOLECULE TYPE: DNA (genomic)	
J u	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
55	(vi)	ORIGINAL SOURCE:	

		(A) ORGANISM: Helicobacter pylori	
5	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 125	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:174:	
	ATGAATTC	AA TTTTTTATTT TGCCA	25
10	(2) INFO	RMATION FOR SEQ ID NO:175:	
15	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
20	(ii)	MOLECULE TYPE: DNA (genomic)	
20	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
25	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
30	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 123	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:175:	
26	AATTCCAT	GG CTATCCAAAT CCG	23
35	(2) INFO	RMATION FOR SEQ ID NO:176:	
40	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 25 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
4.5	(ii)	MOLECULE TYPE: DNA (genomic)	
45	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
50	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
55	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 125	

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	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:176:	
5	ATGAATTCGC CAAAATCGTA GTATT	25
3	(2) INFORMATION FOR SEQ ID NO:177:	
10	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 24 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
15	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
20	(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
25	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 124</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:177:	
30	GATACCATGG AATTTATGAA AAAG	24
50	(2) INFORMATION FOR SEQ ID NO:178:	
35	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 25 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
40	(ii) MOLECULE TYPE: DNA (genomic)	
40	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
45	(vi) ORIGINAL SOURCE:(A) ORGANISM: Helicobacter pylori	
50	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 125</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:178:	
55	TGAATTCGAA AAAGTGTAGT TATAC	25

	(2) INFORMATION FOR SEQ ID NO:179:	
5	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 19 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
10	(ii) MOLECULE TYPE: DNA (genomic)	
10	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
15	(vi) ORIGINAL SOURCE:(A) ORGANISM: Helicobacter pylori	
20	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 119</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:179:	
25	CCCTTCATTT TAGAAATCG	19
	(2) INFORMATION FOR SEQ ID NO:180:	
30	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 20 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
35	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
40	(vi) ORIGINAL SOURCE:(A) ORGANISM: Helicobacter pylori	
45	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 120</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:180:	
50	ATTTCAACCA ATTCAATGCG	20
JU	(2) INFORMATION FOR SEQ ID NO:181:	
55	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 20 base pairs(B) TYPE: nucleic acid	

		(C) STRANDEDNESS: double (D) TOPOLOGY: circular	
_	(ii)	MOLECULE TYPE: DNA (genomic)	
5	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
10	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
15	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 120	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:181:	
20	GCCCCTTT	TG ATTTGAAGCT	20
20	(2) INFO	RMATION FOR SEQ ID NO:182:	•
25	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 22 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
30		MOLECULE TYPE: DNA (genomic)	
		HYPOTHETICAL: NO	
		ANTI-SENSE: NO	
35	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
40	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 122	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:182:	
45	TCGCTCCA	AG ATACCAAGAA GT	22
	(2) INFO	RMATION FOR SEQ ID NO:183:	
50	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 22 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
55	(ii)	MOLECULE TYPE: DNA (genomic)	

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	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
5	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
10	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 122	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:183:	
	CTTGAATT	'AG GGGCAAAGAT CG	22
15	(2) INFO	RMATION FOR SEQ ID NO:184:	
٠	(i)	SEQUENCE CHARACTERISTICS:	
20		(A) LENGTH: 22 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: circular	
25	(ii)	MOLECULE TYPE: DNA (genomic)	
25	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
30	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
35	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 122	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:184:	
40	ATGCGTTT	TT ACCCAAAGAA GT	22
70	(2) INFO	RMATION FOR SEQ ID NO:185:	
45	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 22 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
5 0	(ii)	MOLECULE TYPE: DNA (genomic)	
50	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
55	(\	ORIGINAL SOURCE.	

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		(A) ORGANISM: Helicobacter pylori	
5	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 122	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:185:	
10	ATAACGCC	AC TTCCTTATTG GT	22
10	(2) INFO	RMATION FOR SEQ ID NO:186:	
15	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 19 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
20	(ii)	MOLECULE TYPE: DNA (genomic)	
20	(iii)	HYPOTHETICAL: NO	•
	(iv)	ANTI-SENSE: NO	ı
25	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
30	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 119	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:186:	
35	CTTTGGGT	AA AAACGCATC	19
55	(2) INFO	RMATION FOR SEQ ID NO:187:	
40	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 20 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
45	(ii)	MOLECULE TYPE: DNA (genomic)	
4 J	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
50	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
55	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 120	

	(xi) SEQUENCE DESCRIPTION: SEQ ID N	O:187:
	CGATCTTTGA TCCTAATTCA	20
5	(2) INFORMATION FOR SEQ ID NO:188:	
10	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 19 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
15	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	•
	(iv) ANTI-SENSE: NO	
20	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter py</pre>	/lori
25	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 119</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID 1	IO:188:
30	ATCAAGTTGC CTATGCTGA	19
30	(2) INFORMATION FOR SEQ ID NO:189:	
35	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 22 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
40	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
45	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter page 1</pre>	ylori
50	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 122</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID	NO:189:
	TTGAACACTT TTGATTATGC GG	22

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	(2) INFORMATION FOR SEQ ID NO:190:	
5	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
10	(ii) MOLECULE TYPE: DNA (genomic)	
10	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
15	(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	-
	(ix) FEATURE:	
20	(A) NAME/KEY: misc_feature (B) LOCATION 123	*
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:190:	:
25	GGATTATGCG ATTGTTTTAC AAG	23
23	(2) INFORMATION FOR SEQ ID NO:191:	
30	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 21 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
25	(ii) MOLECULE TYPE: DNA (genomic)	
35	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
40	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	
45	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 121</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:191:	
50	GTCTTTAGCA AAAATGGCGT C	21
J U	(2) INFORMATION FOR SEQ ID NO:192:	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 21 base pairs	
55	(B) TYPE: nucleic acid	

		(C) STRANDEDNESS: double (D) TOPOLOGY: circular	
_	(ii) 1	MOLECULE TYPE: DNA (genomic)	
5	(iii) l	HYPOTHETICAL: NO	
	(iv) i	ANTI-SENSE: NO	
10	(vi) (ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
15		FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 121 SEQUENCE DESCRIPTION: SEQ ID NO:192:	
		A AGAGAGCCTT C	21
20		MATION FOR SEQ ID NO:193:	
25	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 18 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
30		MOLECULE TYPE: DNA (genomic)	
		HYPOTHETICAL: NO	
35		ANTI-SENSE: NO ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
40	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 118	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:193:	
45	CTTATGGGG	G TATTGTCA	18
	(2) INFORMATION FOR SEQ ID NO:194:		
50	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 18 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
55	(ii)	MOLECULE TYPE: DNA (genomic)	

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	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
5	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
10	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 118	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:194:	
15	AGCATGTG	GG TATCCAGC	18
1.5	(2) INFO	RMATION FOR SEQ ID NO:195:	
20	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 19 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	`.
	(ii)	MOLECULE TYPE: DNA (genomic)	•
25	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
30	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
35	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 119	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:195:	
40	AGGTTGTT	GC CTAAAGACT	19
	(2) INFORMATION FOR SEQ ID NO:196:		
45	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 18 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
50	(ii)	MOLECULE TYPE: DNA (genomic)	
	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
c	,		

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		(A) ORGANISM: Helicobacter pylori	
5	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 118	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:196:	
10	CTGCCTCC	AC CTTTGATC	18
	(2) INFO	RMATION FOR SEQ ID NO:197:	
15	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 19 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
20	(ii)	MOLECULE TYPE: DNA (genomic)	
	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
25	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
30	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 119	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:197:	
25	ACCAATAT	CA ATTGGCACT	19
35	(2) INFORMATION FOR SEQ ID NO:198:		
40	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 18 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
	(ii)	MOLECULE TYPE: DNA (genomic)	
45	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
50	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
55	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 1 18	

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	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:198:		
_	ACTTGGAAAA GCTCTGCA	18	
5	(2) INFORMATION FOR SEQ ID NO:199:		
10	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 19 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 		
15	(ii) MOLECULE TYPE: DNA (genomic)		
٠-	(i-i-i-) -HYPOTHETICAL: NO	• =	
	(iv) ANTI-SENSE: NO		
20	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>	*6	
25	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 119</pre>		
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:199:	•	
30	CTTGCTTGTC ATATCTAGC		
30	(2) INFORMATION FOR SEQ ID NO:200:		
35	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 18 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 		
40	(ii) MOLECULE TYPE: DNA (genomic)		
40	(iii) HYPOTHETICAL: NO		
	(iv) ANTI-SENSE: NO		
45	<pre>(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori</pre>		
50	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 118</pre>		
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:200:		
55	GTTGAAGTGT TGGTGCTA	18	

	(2) INFORMATION FOR SEQ ID NO:201:	
5	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 22 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
10	(ii) MOLECULE TYPE: DNA (genomic)	
lU	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
15	(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
20	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 122</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:201:	
25	CAAGCAAGTG GTTTGGTTTT AG	22
23	(2) INFORMATION FOR SEQ ID NO:202:	
30	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 22 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular 	
35	(ii) MOLECULE TYPE: DNA (genomic)	
	(iii) HYPOTHETICAL: NO	
	(iv) ANTI-SENSE: NO	
40	(vi) ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
45	<pre>(ix) FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 122</pre>	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:202:	
50	TGGAAAGAGC AAATCATTGA AG	22
50	(2) INFORMATION FOR SEQ ID NO:203:	
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 21 base pairs	
55	(R) TYPE: nucleic acid	

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		(C) STRANDEDNESS: double (D) TOPOLOGY: circular	
5	(ii)	MOLECULE TYPE: DNA (genomic)	
3	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
10	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
15	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 121	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:203:	
20	GCCCATAA'	TC AAAAAGCCCA T	21
20	(2) INFO	RMATION FOR SEQ ID NO:204:	
25	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 24 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
30		MOLECULE TYPE: DNA (genomic)	
		HYPOTHETICAL: NO	
35	•	ANTI-SENSE: NO ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
40	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 124	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:204:	
45	CTAAAACCA	AA ACCACTTGCT TGTC	24
73	(2) INFOR	RMATION FOR SEQ ID NO:205:	
50	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
55	(ii)	MOLECULE TYPE: DNA (genomic)	

	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
5	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
10	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 116	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:205:	
15	GTAAAACG.	AC GGCCAG	16
13	(2) INFO	RMATION FOR SEQ ID NO:206:	٠
20	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 17 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
25	(ii)	MOLECULE TYPE: DNA (genomic)	
25	(iii)	HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
30	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
35	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 117	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:206:	
40	CAGGAAAC	CAG CTATGAC	17
40	(2) INFO	DRMATION FOR SEQ ID NO:207:	
45	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 21 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	
50	(ii)	MOLECULE TYPE: DNA (genomic)	
30	(iii)) HYPOTHETICAL: NO	
	(iv)	ANTI-SENSE: NO	
55	(vi)	ORIGINAL SOURCE:	

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		(A) ORGANISM: Helicobacter pylori	
5	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 121	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:207:	
10	ATCTTACC	TA TCACCTCAAA T	21
10	(2) INFO	RMATION FOR SEQ ID NO:208:	
<u>15</u>	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 21 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: circular	-
20	(ii)	MOLECULE TYPE: DNA (genomic)	
20	(iii)	HYPOTHETICAL: NO	•
	(iv)	ANTI-SENSE: NO	
25	(vi)	ORIGINAL SOURCE: (A) ORGANISM: Helicobacter pylori	
30	(ix)	FEATURE: (A) NAME/KEY: misc_feature (B) LOCATION 121	
	(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:208:	
	n Ch Ch CCh	A C. A TOTTTOTO A. A.	21

CLAIMS

- An isolated nucleic acid comprising a nucleotide sequence encoding an
 H. pylori polypeptide at least about 60% homologous to an amino acid sequence selected from the group consisting of SEQ ID NO: 74-SEQ ID NO: 146.
 - 2. An isolated nucleic acid comprising a nucleotide sequence encoding an *H. pylori* polypeptide selected from the group consisting of SEQ ID NO: 74-SEQ ID NO: 146.
 - 3. An isolated nucleic acid which encodes an *H. pylori* polypeptide, comprising a nucleotide sequence at least about 60% homologous to a nucleotide sequence selected from the group consisting of SEQ ID NO: 1-SEQ ID NO: 73, or a complement thereof.
 - 4. The isolated nucleic acid of claim 1, comprising a nucleotide sequence selected from the group consisting of SEQ ID NO: 1-SEQ ID NO: 73, or a complement thereof.
 - 5. An isolated nucleic acid molecule encoding an *H. pylori* polypeptide, comprising a nucleotide sequence which hybridizes under stringent hybridization conditions to a nucleic acid molecule comprising the nucleotide sequence selected from the group consisting of SEQ ID NO: 1-SEQ ID NO: 73, or a complement thereof.
 - 6. An isolated nucleic acid comprising a nucleotide sequence of at least 8 nucleotides in length, wherein the sequence hybridizes under stringent hybridization conditions to a nucleic acid having a nucleotide sequence selected from the group consisting of SEQ ID NO: 1-SEQ ID NO: 73, or a complement thereof.
 - 7. An isolated nucleic acid comprising a nucleotide sequence encoding an *H. pylori* cell envelope polypeptide or a fragment thereof, said nucleic acid selected from the group consisting of SEQ ID NO: 3, SEQ ID NO: 25, SEQ ID NO: 48, SEQ ID NO: 16, SEQ ID NO: 10, SEQ ID NO: 45, SEQ ID NO: 35, SEQ ID NO: 37, SEQ ID NO: 7, SEQ ID NO: 39, SEQ ID NO: 55, SEQ ID NO: 18, SEQ ID NO: 19, SEQ ID NO: 28, SEQ ID NO: 30, SEQ ID NO: 52, SEQ ID NO: 54, SEQ ID NO: 56, SEQ ID NO: 58, SEQ ID NO: 1, SEQ ID NO: 42, SEQ ID NO: 14, SEQ ID NO: 43, SEQ ID

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NO: 11, SEQ ID NO: 71, SEQ ID NO: 17, SEQ ID NO: 57, SEQ ID NO: 5, SEQ ID NO: 6, SEQ ID NO: 8, and SEQ ID NO: 21, or a complement thereof.

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- 8. The isolated nucleic acid of claim 7, wherein said *H. pylori* cell envelope polypeptide or a fragment thereof is an *H. pylori* inner membrane polypeptide or a fragment thereof encoded by a nucleic acid selected from the group consisting of SEQ ID NO: 3, SEQ ID NO: 25, and SEQ ID NO: 48, or a complement thereof.
- The isolated nucleic acid of claim 7, wherein said H. pylori cell envelope
 polypeptide or a fragment thereof is an H. pylori outer membrane polypeptide or a fragment thereof encoded by a nucleic acid selected from the group consisting of SEQ ID NO: 16, SEQ ID NO: 10, SEQ ID NO: 45, SEQ ID NO: 35, SEQ ID NO: 37, SEQ ID NO: 7, SEQ ID NO: 39, SEQ ID NO: 55, SEQ ID NO: 18, SEQ ID NO: 19, SEQ ID NO: 28, SEQ ID NO: 30, SEQ ID NO: 52, SEQ ID NO: 54, SEQ ID NO: 56, SEQ ID NO: 58, SEQ ID NO: 1, SEQ ID NO: 42, SEQ ID NO: 14, SEQ ID NO: 43, SEQ ID NO: 11, and SEQ ID NO: 71, or a complement thereof.
 - 10. The isolated nucleic acid of claim 9, wherein said *H. pylori* outer membrane polypeptide or a fragment thereof is an *H. pylori* polypeptide having a terminal phenylalanine residue and a C-terminal tyrosine cluster or a fragment thereof encoded by a nucleic acid selected from the group consisting of SEQ ID NO: 1, SEQ ID NO: 42, SEQ ID NO: 14, SEQ ID NO: 43, SEQ ID NO: 11 and SEQ ID NO:71, or a complement thereof.
- The isolated nucleic acid of claim 9, wherein said H. pylori outer membrane polypeptide or a fragment thereof is an H. pylori polypeptide having a terminal phenylalanine residue or a fragment thereof encoded by a nucleic acid selected from the group consisting of SEQ ID NO: 16, SEQ ID NO: 45, SEQ ID NO: 35, SEQ ID NO: 37, SEQ ID NO: 7, SEQ ID NO: 39, SEQ ID NO: 55, SEQ ID NO: 18, SEQ ID NO: 19, SEQ ID NO: 28, SEQ ID NO: 30, SEQ ID NO: 52, SEQ ID NO: 54, SEQ ID NO: 56, SEQ ID NO: 58, or a complement thereof.
- 12. An isolated nucleic acid comprising a nucleotide sequence encoding an *H. pylori* cell envelope polypeptide or a fragment thereof selected from the group consisting of SEQ ID NO: 76, SEQ ID NO: 98, SEQ ID NO: 121, SEQ ID NO: 89, SEQ ID NO: 83, SEQ ID NO: 118, SEQ ID NO: 108, SEQ ID NO: 110, SEQ ID NO: 80, SEQ ID NO: 112, SEQ ID NO: 128, SEQ ID NO: 91, SEQ ID NO: 92, SEQ ID NO:

101, SEQ ID NO: 103, SEQ ID NO: 125, SEQ ID NO: 127, SEQ ID NO: 129, SEQ ID NO: 131, SEQ ID NO: 74, SEQ ID NO: 115, SEQ ID NO: 87, SEQ ID NO: 116, SEQ ID NO: 84, SEQ ID NO: 144, SEQ ID NO: 90, SEQ ID NO: 130, SEQ ID NO: 78, SEQ ID NO: 79, SEQ ID NO: 81, and SEQ ID NO: 94.

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13. The isolated nucleic acid of claim 12, wherein said *H. pylori* cell envelope polypeptide or a fragment thereof is an *H. pylori* inner membrane polypeptide or a fragment thereof selected from the group consisting of SEQ ID NO: 76, SEQ ID NO: 98, and SEQ ID NO: 121.

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14. The isolated nucleic acid of claim 12, wherein said *H. pylori* cell envelope polypeptide or a fragment thereof is an *H. pylori* outer membrane polypeptide or a fragment thereof selected from the group consisting of SEQ ID NO: 89, SEQ ID NO: 83, SEQ ID NO: 118, SEQ ID NO: 108, SEQ ID NO: 110, SEQ ID NO: 80, SEQ ID NO: 112, SEQ ID NO: 128, SEQ ID NO: 91, SEQ ID NO: 92, SEQ ID NO: 101, SEQ ID NO: 103, SEQ ID NO: 125, SEQ ID NO: 127, SEQ ID NO: 129, SEQ ID NO: 131, SEQ ID NO: 74, SEQ ID NO: 115, SEQ ID NO: 87, SEQ ID NO: 116, SEQ ID NO: 84, SEQ ID NO: 144, SEQ ID NO: 90, and SEQ ID NO: 130.

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15. The isolated nucleic acid of claim 14, wherein said *H. pylori* outer membrane polypeptide or a fragment thereof is an *H. pylori* polypeptide having a terminal phenylalanine residue and a C-terminal tyrosine cluster or a fragment thereof selected from the group consisting of SEQ ID NO: 74, SEQ ID NO: 115, SEQ ID NO: 87, SEQ ID NO: 116, and SEQ ID NO: 84 and SEQ ID NO:144.

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- 16. The isolated nucleic acid of claim 14, wherein said *H. pylori* outer membrane polypeptide or a fragment thereof is an *H. pylori* polypeptide having a terminal phenylalanine residue or a fragment thereof selected from the group consisting of SEQ ID NO: 89, SEQ ID NO: 118, SEQ ID NO: 108, SEQ ID NO: 110, SEQ ID NO: 80, SEQ ID NO: 112, SEQ ID NO: 128, SEQ ID NO: 91, SEQ ID NO: 92, SEQ ID NO: 101, SEQ ID NO: 103, SEQ ID NO: 125, SEQ ID NO: 127, SEQ ID NO: 129, and SEQ ID NO: 131.
- 17. An isolated nucleic acid comprising a nucleotide sequence encoding an
 35 H. pylori secreted polypeptide or a fragment thereof, said nucleic acid selected from the
 group consisting of SEQ ID NO: 72, SEQ ID NO: 32, SEQ ID NO: 51, SEQ ID NO: 2,
 SEQ ID NO: 4, SEQ ID NO: 9, SEQ ID NO: 13, SEQ ID NO: 22, SEQ ID NO: 29, SEQ

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ID NO: 31, SEQ ID NO: 33, SEQ ID NO: 34, SEQ ID NO: 36, SEQ ID NO: 38, SEQ ID NO: 40, SEQ ID NO: 41, SEQ ID NO: 44, SEQ ID NO: 46, SEQ ID NO: 49, SEQ ID NO: 53, SEQ ID NO: 59, SEQ ID NO: 61, SEQ ID NO: 62, SEQ ID NO: 63, SEQ ID NO: 65, SEQ ID NO: 66, SEQ ID NO: 67, and SEQ ID NO: 68, or a complement thereof.

- 18. An isolated nucleic acid comprising a nucleotide sequence encoding an *H. pylori* secreted polypeptide or a fragment thereof selected from the group consisting of SEQ ID NO: 145, SEQ ID NO: 105, SEQ ID NO: 124, SEQ ID NO: 75, SEQ ID NO: 77, SEQ ID NO: 82, SEQ ID NO: 86, SEQ ID NO: 95, SEQ ID NO: 102, SEQ ID NO: 104, SEQ ID NO: 106, SEQ ID NO: 107, SEQ ID NO: 109, SEQ ID NO: 111, SEQ ID NO: 113, SEQ ID NO: 114, SEQ ID NO: 117, SEQ ID NO: 119, SEQ ID NO: 122, SEQ ID NO: 126, SEQ ID NO: 132, SEQ ID NO: 134, SEQ ID NO: 135, SEQ ID NO: 136, SEQ ID NO: 138, SEQ ID NO: 139, SEQ ID NO: 140, and SEQ ID NO: 141.
- 19. An isolated nucleic acid comprising a nucleotide sequence encoding an *H. pylori* cellular polypeptide or a fragment thereof, said nucleic acid selected from the group consisting of SEQ ID NO: 12, SEQ ID NO: 15, SEQ ID NO: 20, SEQ ID NO: 23, SEQ ID NO: 24, SEQ ID NO: 26, SEQ ID NO: 27, SEQ ID NO: 47, SEQ ID NO: 50, SEQ ID NO: 60, SEQ ID NO: 64, SEQ ID NO: 69, SEQ ID NO: 70, and SEQ ID NO: 73, or a complement thereof.
- 20. An isolated nucleic acid comprising a nucleotide sequence encoding an *H. pylori* cellular polypeptide or a fragment thereof selected from the group consisting of SEQ ID NO: 85, SEQ ID NO: 88, SEQ ID NO: 93, SEQ ID NO: 96, SEQ ID NO: 97, SEQ ID NO: 99, SEQ ID NO: 100, SEQ ID NO: 120, SEQ ID NO: 123, SEQ ID NO: 133, SEQ ID NO: 137, SEQ ID NO: 142, SEQ ID NO: 143, and SEQ ID NO: 146.
- 21. A probe comprising a nucleotide sequence consisting of at least 8

 nucleotides of a nucleotide sequence selected from the group consisting of SEQ ID NO:

 1-SEQ ID NO: 73, or a complement thereof.
 - 22. A recombinant expression vector comprising the nucleic acid of any of claims 1, 2, 3, 4, 5, 6, 7, 12, 17, 18, 19 or 20 operably linked to a transcription regulatory element.

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- 23. A cell comprising a recombinant expression vector of claim 22.
- 24. A method for producing an *H. pylori* polypeptide comprising culturing a cell of claim 23 under conditions that permit expression of the polypeptide.
- 25. The method of claim 24, further comprising purifying the polypeptide from the cell.
- 26. A method for detecting the presence of a *Helicobacter* nucleic acid in a sample comprising:
 - (a) contacting a sample with a nucleic acid of any of claims 6 or 21 so that a hybrid can form between the probe and a *Helicobacter* nucleic acid in the sample; and
 - (b) detecting the hybrid formed in step (a), wherein detection of a hybrid indicates the presence of a *Helicobacter* nucleic acid in the sample.
 - 27. An isolated *H. pylori* polypeptide comprising an amino acid sequence at least about 60% homologous to an *H. pylori* polypeptide selected from the group consisting of SEQ ID NO: 74-SEQ ID NO: 146.
 - 28. An isolated *H. pylori* polypeptide which is encoded by a nucleic acid comprising a nucleotide sequence at least about 60% homologous to a nucleotide sequence selected from the group consisting of SEQ ID NO: 1-SEQ ID NO: 73.
- 25 29. The isolated *H. pylori* polypeptide of claim 28, wherein said polypeptide is encoded by a nucleotide sequence selected from the group consisting of SEQ ID NO: 1-SEO ID NO: 73.
- 30. An isolated *H. pylori* polypeptide which is encoded by a nucleic acid which hybridizes under stringent hybridization conditions to a nucleic acid selected from the group consisting of SEQ ID NO: 1-SEQ ID NO: 73, or a complement thereof.
 - 31. An isolated *H. pylori* polypeptide comprising an amino acid sequence selected from the group consisting of SEQ ID NO: 74-SEQ ID NO: 146.
 - 32. An isolated *H. pylori* cell envelope polypeptide or a fragment thereof, wherein said polypeptide is selected from the group consisting of SEQ ID NO: 76, SEQ

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ID NO: 98, SEQ ID NO: 121, SEQ ID NO: 89, SEQ ID NO: 83, SEQ ID NO: 118, SEQ ID NO: 108, SEQ ID NO: 110, SEQ ID NO: 80, SEQ ID NO: 112, SEQ ID NO: 128, SEQ ID NO: 91, SEQ ID NO: 92, SEQ ID NO: 101, SEQ ID NO: 103, SEQ ID NO: 125, SEQ ID NO: 127, SEQ ID NO: 129, SEQ ID NO: 131, SEQ ID NO: 74, SEQ ID NO: 115, SEQ ID NO: 87, SEQ ID NO: 116, SEQ ID NO: 84, SEQ ID NO: 144, SEQ ID NO: 90, SEQ ID NO: 130, SEQ ID NO: 78, SEQ ID NO: 79, SEQ ID NO: 81, and SEQ ID NO: 94.

- 33. The isolated polypeptide of claim 32, wherein said *H. pylori* cell

 envelope polypeptide or a fragment thereof is an *H. pylori*-inner-membrane-polypeptide or a fragment thereof selected from the group consisting of SEQ ID NO: 76, SEQ ID NO: 98, and SEQ ID NO: 121.
- 34. The isolated polypeptide of claim 32, wherein said *H. pylori* cell envelope polypeptide or a fragment thereof is an *H. pylori* outer membrane polypeptide or a fragment thereof selected from the group consisting of SEQ ID NO: 89, SEQ ID NO: 83, SEQ ID NO: 118, SEQ ID NO: 108, SEQ ID NO: 110, SEQ ID NO: 80, SEQ ID NO: 112, SEQ ID NO: 128, SEQ ID NO: 91, SEQ ID NO: 92, SEQ ID NO: 101, SEQ ID NO: 103, SEQ ID NO: 125, SEQ ID NO: 127, SEQ ID NO: 129, SEQ ID NO: 131, SEQ ID NO: 74, SEQ ID NO: 115, SEQ ID NO: 87, SEQ ID NO: 116, SEQ ID NO: 84, SEQ ID NO: 144, SEQ ID NO: 90, and SEQ ID NO: 130.
 - 35. The isolated polypeptide of claim 34, wherein said *H. pylori* outer membrane polypeptide or a fragment thereof is an *H. pylori* polypeptide having a terminal phenylalanine residue and a C-terminal tyrosine cluster or a fragment thereof selected from the group consisting of SEQ ID NO: 74, SEQ ID NO: 115, SEQ ID NO: 87, SEQ ID NO: 116, and SEQ ID NO: 84 and SEQ ID NO:144.
- 36. The isolated polypeptide of claim 34, wherein said *H. pylori* outer
 30 membrane polypeptide or a fragment thereof is an *H. pylori* polypeptide having a
 terminal phenylalanine residue or a fragment thereof selected from the group consisting
 of SEQ ID NO: 89, SEQ ID NO: 118, SEQ ID NO: 108, SEQ ID NO: 110, SEQ ID NO:
 80, SEQ ID NO: 112, SEQ ID NO: 128, SEQ ID NO: 91, SEQ ID NO: 92, SEQ ID NO:
 101, SEQ ID NO: 103, SEQ ID NO: 125, SEQ ID NO: 127, SEQ ID NO: 129, and SEQ
 35 ID NO: 131.

An isolated *H. pylori* cell envelope polypeptide or a fragment thereof, wherein said polypeptide is encoded by a nucleic acid selected from the group consisting of SEQ ID NO: 3, SEQ ID NO: 25, SEQ ID NO: 48, SEQ ID NO: 16, SEQ ID NO: 10, SEQ ID NO: 45, SEQ ID NO: 35, SEQ ID NO: 37, SEQ ID NO: 7, SEQ ID NO: 39, SEQ ID NO: 55, SEQ ID NO: 18, SEQ ID NO: 19, SEQ ID NO: 28, SEQ ID NO: 30, SEQ ID NO: 52, SEQ ID NO: 54, SEQ ID NO: 56, SEQ ID NO: 58, SEQ ID NO: 1, SEQ ID NO: 42, SEQ ID NO: 14, SEQ ID NO: 43, SEQ ID NO: 11, SEQ ID NO: 71, SEQ ID NO: 17, SEQ ID NO: 57, SEQ ID NO: 5, SEQ ID NO: 6, SEQ ID NO: 8, and SEQ ID NO: 21.

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38. The isolated polypeptide of claim 37, wherein said *H. pylori* cell envelope polypeptide or a fragment thereof is an *H. pylori* inner membrane polypeptide or a fragment thereof encoded by a nucleic acid selected from the group consisting of SEQ ID NO: 3, SEQ ID NO: 25, and SEQ ID NO: 48.

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and the isolated polypeptide of claim 37, wherein said *H. pylori* cell envelope polypeptide or a fragment thereof is an *H. pylori* outer membrane polypeptide or a fragment thereof encoded by a nucleic acid selected from the group consisting of SEQ ID NO: 16, SEQ ID NO: 10, SEQ ID NO: 45, SEQ ID NO: 35, SEQ ID NO: 37, SEQ ID NO: 7, SEQ ID NO: 39, SEQ ID NO: 55, SEQ ID NO: 18, SEQ ID NO: 19, SEQ ID NO: 28, SEQ ID NO: 30, SEQ ID NO: 52, SEQ ID NO: 54, SEQ ID NO: 56, SEQ ID NO: 58, SEQ ID NO: 1, SEQ ID NO: 42, SEQ ID NO: 14, SEQ ID NO: 43, SEQ ID NO: 11, and SEQ ID NO: 71.

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40. The isolated polypeptide of claim 39, wherein said *H. pylori* outer membrane polypeptide or a fragment thereof is an *H. pylori* polypeptide having a terminal phenylalanine residue and a C-terminal tyrosine cluster or a fragment thereof encoded by a nucleic acid selected from the group consisting of SEQ ID NO: 1, SEQ ID NO: 42, SEQ ID NO: 14, SEQ ID NO: 43, SEQ ID NO: 11 and SEQ ID NO:71.

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41. The isolated polypeptide of claim 39, wherein said *H. pylori* outer membrane polypeptide or a fragment thereof is an *H. pylori* polypeptide having a terminal phenylalanine residue or a fragment thereof encoded by a nucleic acid selected from the group consisting of SEQ ID NO: 16, SEQ ID NO: 45, SEQ ID NO: 35, SEQ ID NO: 37, SEQ ID NO: 7, SEQ ID NO: 39, SEQ ID NO: 55, SEQ ID NO: 18, SEQ ID NO: 19, SEQ ID NO: 28, SEQ ID NO: 30, SEQ ID NO: 52, SEQ ID NO: 54, SEQ ID NO: 56, SEQ ID NO: 58.

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- 42. An isolated *H. pylori* cellular polypeptide or a fragment thereof, wherein said polypeptide is selected from the group consisting of SEQ ID NO: 85, SEQ ID NO: 88, SEQ ID NO: 93, SEQ ID NO: 96, SEQ ID NO: 97, SEQ ID NO: 99, SEQ ID NO: 100, SEQ ID NO: 120, SEQ ID NO: 123, SEQ ID NO: 133, SEQ ID NO: 137, SEQ ID NO: 142, SEQ ID NO: 143, and SEQ ID NO: 146.
- 43. An isolated *H. pylori* cellular polypeptide or a fragment thereof, wherein said polypeptide is encoded by a nucleic acid selected from the group consisting of SEQ ID NO: 12, SEQ ID NO: 15, SEQ ID NO: 20, SEQ ID NO: 23, SEQ ID NO: 24, SEQ ID NO: 26, SEQ ID NO: 27, SEQ ID NO: 47, SEQ ID NO: 50, SEQ ID NO: 60, SEQ ID NO: 64, SEQ ID NO: 69, SEQ ID NO: 70, and SEQ ID NO: 73.
- 44. An isolated *H. pylori* secreted polypeptide or a fragment thereof, wherein said polypeptide is selected from the group consisting of SEQ ID NO: 145, SEQ ID NO: 105, SEQ ID NO: 124, SEQ ID NO: 75, SEQ ID NO: 77, SEQ ID NO: 82, SEQ ID NO: 86, SEQ ID NO: 95, SEQ ID NO: 102, SEQ ID NO: 104, SEQ ID NO: 106, SEQ ID NO: 107, SEQ ID NO: 109, SEQ ID NO: 111, SEQ ID NO: 113, SEQ ID NO: 114, SEQ ID NO: 117, SEQ ID NO: 119, SEQ ID NO: 122, SEQ ID NO: 126, SEQ ID NO: 132, SEQ ID NO: 134, SEQ ID NO: 135, SEQ ID NO: 136, SEQ ID NO: 138, SEQ ID NO: 139, SEQ ID NO: 140, and SEQ ID NO: 141.
 - 45. An isolated *H. pylori* secreted polypeptide or a fragment thereof, wherein said polypeptide is encoded by a nucleic acid selected from the group consisting of SEQ ID NO: 72, SEQ ID NO: 32, SEQ ID NO: 51, SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 9, SEQ ID NO: 13, SEQ ID NO: 22, SEQ ID NO: 29, SEQ ID NO: 31, SEQ ID NO: 33, SEQ ID NO: 34, SEQ ID NO: 36, SEQ ID NO: 38, SEQ ID NO: 40, SEQ ID NO: 41, SEQ ID NO: 44, SEQ ID NO: 46, SEQ ID NO: 49, SEQ ID NO: 53, SEQ ID NO: 59, SEQ ID NO: 61, SEQ ID NO: 62, SEQ ID NO: 63, SEQ ID NO: 65, SEQ ID NO: 66, SEQ ID NO: 67, and SEQ ID NO: 68.
 - 46. A fusion protein comprising an *H. pylori* polypeptide which comprises an amino acid sequence selected from the group consisting of SEQ ID NO: 74-SEQ ID NO: 146 operatively linked to a non-*H. pylori* polypeptide.

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- 47. A vaccine formulation for prophylaxis or treatment of an *H. pylori* infection comprising an effective amount of at least one isolated nucleic acid of any of claims 1, 2, 3, 4, 5, 6, 7, 12, 17, 18, 19, or 20.
- 48. A vaccine formulation for prophylaxis or treatment of an *H. pylori* infection comprising an effective amount of at least one *H. pylori* polypeptide or a fragment thereof of any of claims 26, 27, 28, 29, 30, 31, 32, 37, 42, 43, 44 or 45.
- 49. A vaccine formulation of claim 47, further comprising a pharmaceutically acceptable carrier.
 - 50. A vaccine formulation of claim 48, further comprising a pharmaceutically acceptable carrier.
- 15 51. A vaccine formulation of claim 49, wherein the pharmaceutically acceptable carrier comprises an adjuvant.
 - 52. A vaccine formulation of claim 50, wherein the pharmaceutically acceptable carrier comprises an adjuvant.
 - 53. A vaccine formulation of claim 49, wherein the pharmaceutically acceptable carrier comprises a delivery system.
- 54. A vaccine formulation of claim 50, wherein the pharmaceutically acceptable carrier comprises a delivery system.
 - 55. A vaccine formulation of claim 53, wherein the delivery system comprises a live vector.
- 30 56. A vaccine formulation of claim 54, wherein the delivery system comprises a live vector.
 - 57. A vaccine formulation of claim 55, wherein the live vector is a bacteria or a virus.
 - 58. A vaccine formulation of claim 56, wherein the live vector is a bacteria or a virus.

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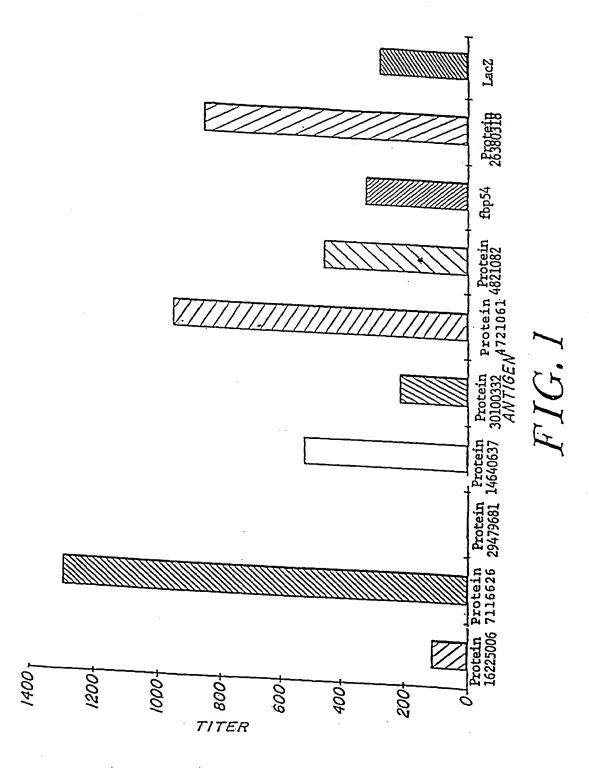
- 59. A vaccine formulation of claim 53, wherein the pharmaceutically acceptable carrier further comprises an adjuvant.
- 5 60. A vaccine formulation of claim 54, wherein the pharmaceutically acceptable carrier further comprises an adjuvant.

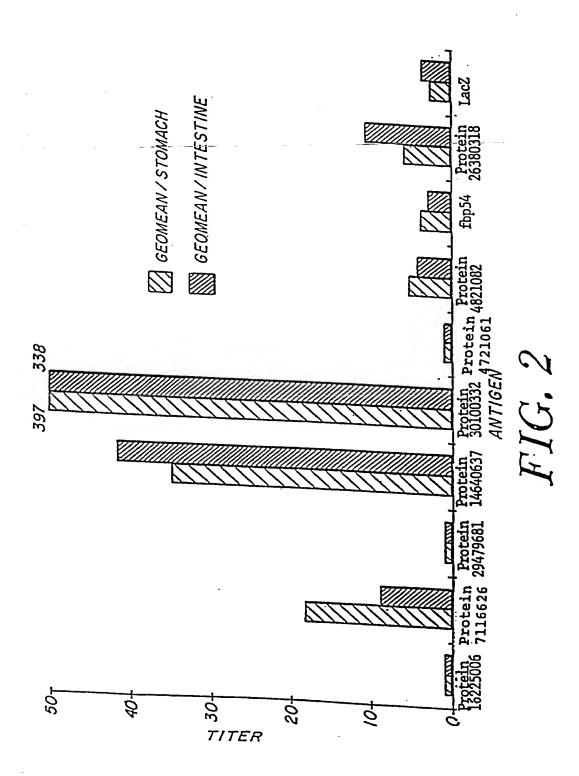
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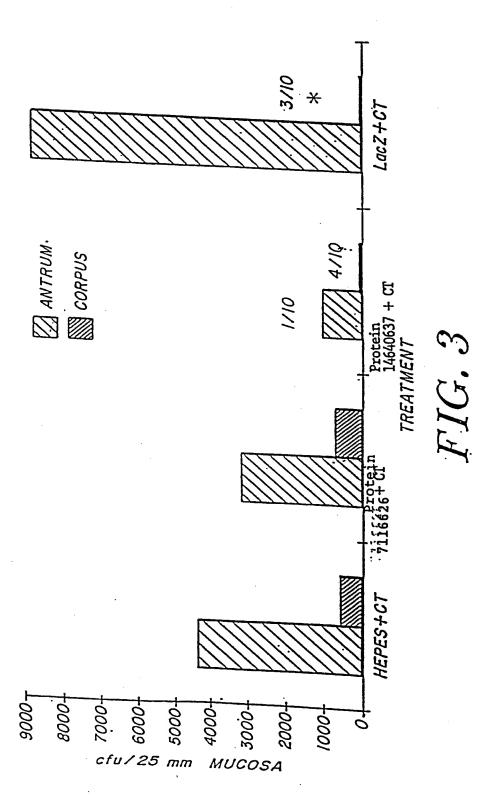
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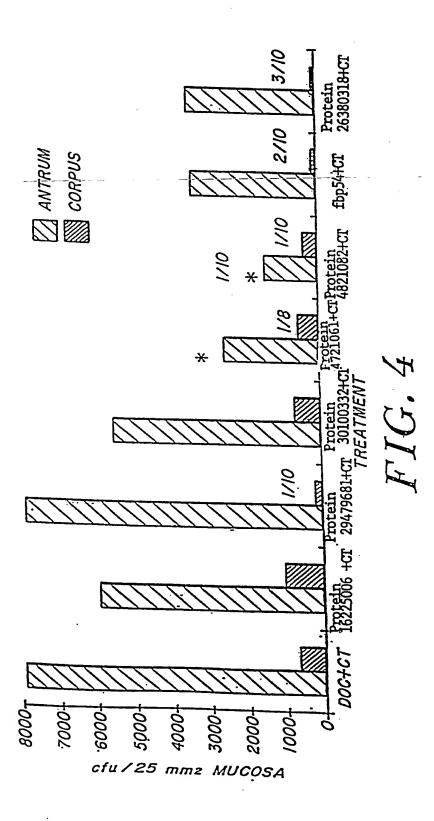
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- 61. A method of treating or reducing a risk of *H. pylori* infection in a subject comprising administering to a subject a vaccine formulation of claim 47, such that treatment or reduction of risk of *H. pylori* infection occurs.
 - 62. A method of treating or reducing a risk of *H. pylori* infection in a subject comprising administering to a subject a vaccine formulation of claim 48, such that treatment or reduction of risk of *H. pylori* infection occurs.
- 63. A method of producing a vaccine formulation comprising: combining at least one isolated *H. pylori* polypeptide or a fragment thereof selected from the group consisting of SEQ ID NO: 74-SEQ ID NO: 146 with a pharmaceutically acceptable carrier to thereby form a vaccine formulation.
 - 64. A method of producing a vaccine formulation comprising:
- (a) providing at least one isolated *H. pylori* polypeptide or a fragment thereof selected from the group consisting of SEQ ID NO: 74-SEQ ID NO: 146; and
- (b) combining at least one said isolated *H. pylori* polypeptide or a fragment thereof with a pharmaceutically acceptable carrier to thereby form a vaccine formulation.
 - 65. A method of producing a vaccine formulation comprising:
- (a) culturing a cell under condition that permit expression of an H.
 30 pylori polypeptide or a fragment thereof selected from the group consisting of SEQ ID NO: 74-SEQ ID NO: 146;
 - (b) isolating said *H. pylori* polypetide from said cell; and
 - (c) combining at least one said isolated *H. pylori* polypeptide or a fragment thereof with a pharmaceutically acceptable carrier to thereby form a vaccine formulation.









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aa Se	_	BLOCK A
74		MIKRIAC-ILSLSASLALAGEVNGFFMGAGYQQGRYGPYNSNY
115		MIKRIAC-ILSLSASLALAGEVNGFFMGAGYOOGRYGPYNSNY
87		MKKFFSQSLLAL-IISMNAVSGMDGNGVFLGAGYLQGDAQMHADIN
116		MKKFFSQSLLAL-IISMNAVSGMDGNGVFLGAGYLQGDAQMHADIN
84		MARULMKKFVALGLLSAVLSSSLLAEGIGVYIGTNYQLGDARLNSNIYNTGDCTGS
		* . * . *
		BLOCK B BLOCK C
74		KWFGARV
115		KWFGARV
87		KHFGLRL
116		KHFGLRL
84	VVC	GCPPGLTANKHNPGGTNINWHSKYANGALTGFGLNVGYKKFFQFKSLDMTSKWFGFRV
		. * . * . * . * . * * * * .
		· · · · · · · · · · · · · · · · · · ·
74	YGF	FLDWFNTSGTEHTKTNLLTYGGGGD
115	YGF	FLIWFNTSGTEHTKTNLLTYGGGGD
87	YGF	FLYAHANSIKLKNPNYNSEAAQVASQILGKQEINRLTNIADPRTFEPNMLTYGGAMD
116	YGF	FIYAHANSIKLKNPNYNSEAAQVASQILGKQEINRLTNIADPRTFEPNMLTYGGAMD
84	YGI	FTYGHADLGKQVYAPNKIQLDMVSWGVGSD
	**	<u> </u>
		BLOCK D
74	LIV	NLIPLDKFALGLIGGVQLAGNTWMFPYDVNQ
115		NLIPLDKFALGLIGGVQLAGNTWMFPYDVNQ
87	VMV	NVINNGIMSLGAFGGIQLAGNSWLMATPSFEGILVEQALV
116	VMV	nvinngimsigafggiqlagnswlmatpsfegilveqalv
84	LLA	DIIDKDNASFGIFGGVAIGGNTWKSSAANYWKEQIIEAKGPDVCTPTYCNPNAPYST
	• •	·* · * **. · **.*
		BLOCK F
7.4		-TRFQFLWNLGGRMRVGDRSAFEAGVKFPMVNQGSKDVGLTRYYSWYV
115		-TRFQFLWNLGGRMRVGDRSAFEAGVKFPMVNQGSKDVGLTRYYSWYV
87	SKK	ATSFQFLFNVGARLRILKHSSIEAGVKFPMLKKNPYITAKNLDIGFRRVYSWYV
116	SKK	ATSFQFLFNVGARLRILKHSSIEAGVKFPMLKKNPYITAKNLDIGFRRVYSWYV
84	NTS	TV4FQVWLNFGVRANIYKHNGVEFGVRVPLLINKFLSAGPNATNLYYHLKRDYSLYL
		** * * * * * **. *
		——————————————————————————————————————
74		FTF
115		FTF
87	NYV	FTF
116	NYV	FTF
84		YTF
	*	<u>·**</u>

FIGURE 5

aaSeqID#	
83 89	MRKLFIPLLLFSALEANEKNGFFIEAGFETGLLEGTQTQEKRHTTTKNTYATYNYLPTDT
108	MRKLFIPLLLFSALEANEKNGFFIEAGFETGLLEGTQTQEKRHTTTKNTYATYNYLPTDT
118	MRKLFIPLLLFSALEANEKNGFFIEAGFETGLLEGTQTQEKRHTTTKNTYATYNYLPTDT

83	ILKRAANLFTNAEAISKLKFSSLSPVRVLYMYNGQLTIENFLPYNLMNVKLSFTDAQGNV
89	
108	ILKRAANLFTNAEAISKLKFSSLSPVRVLYMYNGQLTIENFLPYNLNNVKLSFTDAQGNT
118	ILKRAANLFTNAEAISKLKFSSLSPVRVLYMYNGQLTIENFLPYNLNNVKLSFTDAQGNV
83	IDLGVIETIPKHSKIVLPGEAFDSLKIDPYTLFLPKIEATSTSISDANTORVFET
89	VIETIPKHSKIVLPGEAFDSLKEAFDKIDPYTFFFPKFEATSTSISDTNTQRVFET
108	IDLGVIETIPKHSKIVLPGEAFDSLKEAFDKIDPYTLPLPKFEATSTSISDINTQRVFET
118	IDLGVIETIPKHSKIVLPGEAFDSLKIDPYTLFLPKIEATSTSISDANTQRVFET

83	LNKIKTNLVVNYRNENKFKDHENHWEAFTPQTAEEFTNLMLNMIAVLDS
89	LNNIKTNLIMKYSNENPNNFNTCPYNNNGNTKNDCWQNFTPQTAEEFTNLMLNMIAVLDS
108	LNNIKTNLIMKYSNENPNNFNTCPYNNNGNTKNDCWQNFTPQTAEEFTNLMLNMIAVLDS
118	LNKIKTNLVVNYRNENKFKDHENHWEAFTPQTAEEFTNLMLNMIAVLDS
83	QSWGDAILNAPFEFTNSPTDCDNDPSKCVNPGTNGLVNSKVDQKYVLNKQDIVNKFKNKA
89	QSWGDAILNAPFEFTNSSTDCDSDPSKCVNPGVNGRVDTKVDQQYILNKQGIINNFRKKI
108	QSWGDAILNAPFEFTNSSTDCDSDPSKCVNPGVNGRVDTKVDQQYILNKQGIINNFRKKI
118	QSWGDAILNAPFEFTNSPTDCDNDPSKCVNPGTNGLVNSKVDQKYVLNKQDIVNKFKNKA

83	DLDVIVLKDSGVVGLGSDITPSNNDDGKHYGQLGVVASALDPKKLFGDNLKTINLEDLRT
89	EIDAVVLKNSGVVGLANGYGNDG-EYGTLGVEAYALDPKKLFGNDLKTINLEDLRT
108	EIDAVVLKNSGVVGLANGYGNDG-EYGTLGVEAYALDPKKLFGNDLKTINLEDLRT
118	DLDVIVLKDSGVVGLGSDITPSNNDDGKHYGOLGVVASALDPKKLFGDNLKTINLEDLRT
:	。 · · · · · · · · · · · · · · · · · · ·
83	ILHEFSHTKGYGHNGNMTYORVPVTKDGOVEKDSNGKPKDSDGLPYNVC
89	ILHEFSHTKGYGHNGNMTYQRVPVTKDGQVEKDSNGKPKDSDGLPYNVCSLYGGSNQPAF
108	ILHEFSHTKGYGHNGNMTYQRVPVTKDGQVEKDSNGKPKDSDGLPYNVCSLYGGSNQPAF
118	ILHEFSHTKGYGHNGNMTYQRVPVTKDGQVEKDSNGKPKDSDGLPYNVCSLYGGSNQPAF
113	**************************************
83	
89	PSNYPNSIYHNCADVPAGFLGVTAAVWQQLINQNALPINYANLGSQTNYNLNASLNTQDI
108	PSNYPNSIYHNCADVPAGFLGVTAAVWQQLINQNALPINYANLGSOTNYNLNASLNTODL
118	PSNYPNSIYHNCADVPAGFLGVTAAVWQQLINQNALPINYANLGSQTNYNLNASLNTQDI

FIGURE 6 (Cont'd)

83 83	ANSMLSTIQKTFVTSSVTNHHFSNASQSFRSPILGVNAKIGYQNYFNDFIGLAYYGIIKY	
108 118	ANSMLSTIQKTFVTSSVTNHHFSNASQSFRSPILGVNAKIGYQNYFNDFIGLAYYGIIKY ANSMLSTIQKTFVTSSVTNHHFSNASQSFRSPILGVNAKIGYQNYFNDFIGLAYYGIIKY	
83	**************************************	
89	NYAKAVNQKVQQLSYGGGIDLLLDFITTYSNKNSPTGIQTKRNF5SSFGIFGGLRGLYNS	
108	NYAKAVAQKVQQLSXGGGIDLLLDFITTXSNKNSPTGIQTKRNFSSSFGIFGGLRGLXNS	
118	NYAKAVNQKVQQLSYGGGIDLLLDFITTYSNKNSPTGIQTKRNFSSSFGIFGGLRGLYNS ************************************	
83		
89	YYVLNKVKGSGNLDVATGLNYRYKIISKY <i>S</i> VGISIPLIQRKASVVSSGGDYTNSFVFNEGA	
108	YYVLANKVKGSGNLDVATGLAYRYKHSKYSVGISIPLIQRKASVVSSGGDYTNSFVFNEGA	
118	YYVLNKVKGSGNLDVATGLNYRYKHSKYSVGISIPLIQRKASVVSSGGDYTNSFVFNEGA	
1		
83		
83	SHFKVFFNYGGCF	
108	SHFKVFFNYGWVF	
118	SHFKVFFNYGWVF	

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aaSeqID 80 112	BLOCK A VLKFQKLPLLFVSILYNQSPLLAFDYKFSGVAESVSKVGFNHSKLNSKEGIFPTATFVTARIGVIFSSYATMSGLPSSGTLNSW . * * * * *
80	BLOCK B TIKLQVDSNLLPKNIEKHSLKIGVGGILGALAYDSTKTLIDQATHQIYGSELFYLIGRWW
112	NGLGGNVRNTKVYGKFAYHHLQKYLLIDLIARFK
80	GFI GNA PWKDSI TESDA HTDARWI VAICAT THE COLOR
112	GFLGNAPWKDSLIESDAHTRNYVLYNSYLFYSYGDKFHLKLGRYLSNMDFMSSYTQGFEL TQGGYIFRYNTDDYLPLNSTFYMGGVTTVRGFRNG
	* * * * * * * * * * * * * * * * * * *
	BLOCK C
80	DYKINSKIALKWFSSFGRALAFGQWIRDWYAPIVTEDGRKEVYDGIHAAQLYFSKHVQV
112	SITPKDEFGLWLG
	* ** *. * * ** *
80	MPFAYFSPKIYGAPGVKIHIDSNPKFKGLGLRAQTTINVIFPVYAKDLYDVYWRNSKIGE
112	YGVLKAAKMRLAWFFDFGFLTFKTPTRGSFFYN
	**
	BLOCK D
80	WGASLLIHQREDYNEFNFGFGYYQNFGNANARIGWYGNPIPFNYRNNSVYGGVFSNAITA
112	APTTTANFX DYGVVGAGFERATWRASTGLQIEWISPMGPLVL
	* * * * *
80	DAVSGYVFGGGVYRGFLWGILGRYTYATRASERSINLNLGYKWGSFARVDVNLEYYVVSM
112	OWG
	·* · **
	BLOCK E
80	HNGYRLDYLTGPFNKAFKADAQDRSNLMVSMKFFF
112	GNGKKCKGLCFNPNMNDYTQHFEFSMGTRF
	** . *. ** .* . **

FIGURE 7

aa SeqID#

---EVALKLNYHPASE HVTPLDFNYPVHIVQAPQNHHVVGILMPRIQVSDN-LKPYIDKFQDALINQIQTIFEKRG **ILLLRPAFQYSDNIAKEYENKFKNQTTLKVEEILQNQG** YQVLRFQ--DEKALNVQJKKKIFSVLDLKGWVGILEDLKMNLKDPNSP}-NLDTLVDQSS YKVINVDSSDKDDFSFA¶KKEGYLAVAMNGEIVLRPDPKRTIQKKSEPβLLFSTGLDKME **SSVWFNFYEPESNRVVHDFAVEV**bTFQALTYTYTSTNNASGGFNSSKSVIHENL RVLIPAGFVKVTILEPMSGESLDSFTMDLBELDIQEKFLKTTHSSHSGG--LVSTMVKGT MGCSFIFKKVRVYSKMLVALGLSSVLIGCAMNPSAETKKPNDAKNQQPVQTHERMTTSSE DKNREDAIHKILNRMYAVVMKKAVTILTKENIAKYRDAIDRMKGFKSSMPQKK D-NSNDAIKSALNKIFASIMQEMDK||UTQRNLESYQKDAKELKNKR|NR----MKTNGHFKDF-AWKKCFLGASVVALLVGCSPHIIETN-----BLOCK E BLOCK A BLOCK B BLOCK C BLOCK D KVQALDEK--11111 130 81 130 81 81 130 81 130 81

FIGURE 8

INTERNATIONAL SEARCH REPORT

International application No. PCT/US97/19575

A. CLASSIFICATION OF SUBJECT MATTER IPC(6) :A01N 43/04; A61K 31/70; C12Q 1/68					
US CL:514/44; 435/6 According to International Patent Classification (IPC) or to both national classification and IPC					
B. FIELDS SEARCHED	ileation (1. c) or to both	i national onesmication and it			
Minimum documentation searched (class	ification system followe	ed by classification symbols)			
U.S. : 514/44; 435/6					
Documentation searched other than minin GENEBANK	num documentation to the	e extent that such documents are included	in the fields searched		
Electronic data base consulted during the NONE	e international search (n	ame of data base and, where practicable	e, search terms used)		
C. DOCUMENTS CONSIDERED	TO BE RELEVANT				
Category* Citation of document, v	with indication, where a	ppropriate, of the relevant passages	Relevant to claim No.		
and Demonstration	of Diversity at the ober 1992, Vol. 17	icobacter pylori Genome Map e Genome Level. Journal of 74, No. 21, pages 6800-6806,	1-65		
Helicobacter pylori	detected by PCR-rch. 1992, Vol. 2	among clinical isolates of based RAPD fingerprinting. 0, No. 19, pages 5137-5142,	1-65		
Further documents are listed in the	e continuation of Box C	See patent family annex.			
"A" document defining the general state of the to be of particular relevance	e art which is not considered	"T" later document published after the inte date and not in conflict with the appli the principle or theory underlying the	cation but cited to understand		
*E" carlier document published on or after th		"X" document of particular relevance, the considered novel or cannot be consider when the document is taken alone	claimed invention cannot be ed to involve an inventive step		
 document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) document referring to an oral disclosure, use, exhibition or other means 		"Y" document of particular relevance, the considered to involve an inventive combined with one or more other such being obvious to a person skilled in the	step when the document is documents, such combination		
PP document published prior to the international filing data but later than the priority data claimed		•			
Date of the actual completion of the inter	national search	Date of mailing of the international search report			
27 FEBRUARY 1998		1 3 MAR 1998			
Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231 Facsimile No. (703) 305-3230 Authorized officer GINNY PORTNER Telephone No. (703) 308-0196					

INTERNATIONAL SEARCH REPORT

International application No. PCT/US97/19575

Box 1 Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This international report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:
2. Claims Nos.: because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:
Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box 11 Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This International Searching Authority found multiple inventions in this international application, as follows:
Please See Extra Sheet.
1. As all required additional search fees were timely paid by the applicant, this international search report covers all searchable
As all required additional search rees were timely paid by the application, and claims.
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
As only some of the required additional search fees were timely paid by the applicant, this international search report covers
only those claims for which fees were paid, specifically claims Nos.: 1-65, SEQ. ID Nos. 1, 7, 8, 11, 37, 39, 43, 45, 55, 61, 74, 80, 81 and 112
4. No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark on Protest The additional search fees were accompanied by the applicant's protest.
X No protest accompanied the payment of additional search fees.

INTERNATIONAL SEARCH REPORT

International application No. PCT/US97/19575

BOX II. OBSERVATIONS WHERE UNITY OF INVENTION WAS LACKING This ISA found multiple inventions as follows:

This application contains the following inventions or groups of inventions which are not so linked as to form a single inventive concept under PCT Rule 13.1. In order for all inventions to be searched, the appropriate additional search fees must be paid.

Group I, claim(s) 1-26, 47, 49, 51, 53, 55, 57, 59, and 61, drawn to no fewer than 135 nucleic acid molecules, vectors containing the nucleic acid molecules, DNA encoding fragments of the polypeptides encoded by the no fewer than 135 different DNAs, organism transformed with the nucleic acid molecules, vaccines and methods of producing polypeptides encoded by the no fewer than 135 different nucleic acid molecules.

Group II, claim(s) 27-46, 48, 50, 52, 54, 56, 58, 60, and 62-65 are, drawn to no fewer than 73 polypeptides encoded by a subset of the encoding DNA mentioned in Group I.

This application contains claims directed to more than one species of the generic invention. These species are deemed to lack Unity of Invention because they are not so linked as to form a single inventive concept under PCT Rule 13.1. In order for more than one species to be searched, the appropriate additional search fees must be paid. The species are as follows:

Group I contains a separate DNA species for each sequence mentioned. Therefore, there is a minimum of 135 species.

Group II contains at least one polypeptide for each DNA sequence mentioned. Therefore, this is a minimum of 73 species in Group II.

For either Group that applicant elects, a total of 10 (ten) specified sequences will be searched and no more than 4 (four) specified sequences will be searched for each additional fee paid; if no additional fee is paid and no election indicated the first 10 sequences appearing in Group I will be searched.

and it considers that the International Application does not comply with the requirements of unity of invention (Rules 13.1, 13.2 and 13.3) for the reasons indicated below:

The inventions listed as Groups I and II do not relate to a single inventive concept under PCT Rule 13.1 because, under PCT Rule 13.2, they lack the same or corresponding special technical features for the following reasons: The polypeptide encoding DNAs, vectors containing them, organisms transformed with them and methods of polypeptide production using them of Group I are materially different from each other and are therefore independent from the polypeptides of Group II. Additionally, none of the products or methods of Group I is needed to make the polypeptides of Group II.

The species listed above do not relate to a single inventive concept under PCT Rule 13.1 because, under PCT Rule 13.2, the species lack the same or corresponding special technical features for the following reasons: There is no relationship between or among the various nucleotide and amino acid sequences mentioned in the claims.

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08/739,150

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WORLD INTELLECTUAL PROPERTY ORGANIZATION International Bureau



INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)								
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(63) Related by Continuation (CON) or Continuation-in-Part (CIP) to Earlier Applications

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(72) Inventors; and

(75) Invent rs/Applicants (for US only): SMITH, Douglas [US/US]; 2 Mayflower Lane, Gloucester, MA 01930 (US). (81) Designated States: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, EE, ES, FI, GB, GE, GH, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, ARIPO patent (GH, KE, LS, MW, SD, SZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG).

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(57) Abstract

Recombinant or substantially pure preparations of H. pylori polypeptides are described. The nucleic acids encoding the polypeptides also are described. The H. pylori polypeptides are useful for diagnostics and vaccine compositions.

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